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Susan ITailly Access DB# SEARCH REQUEST FORM Scientific and Technical Inf rmati n Center tequester's Full Name: Christino Maugla Examiner #: 79006, Date: Augus Int Unit: 1637 Phone Number 30 8 - 3617 Serial Number: 09/829 Results Format Preferred (circle): PAPER A Box and Bldg/Room Location: 10A/6 f more than one search is submitted, please prioritize searches in order of need. Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract. Title of Invention: Hiroshia Inventors (please provide full names): Technical Info. Specialist CM1 6805 Tel: 305-4053 Earliest Priority Filing Date: \*For Sequence Searches Only\* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the (Structures) are the Same. appropriate serial number. lains 1-8 are more specific, I expect some for markush groups to surface in old interatine. but look for one attached to a nucleic Acid or some (60) Aructure Specification for examples 1-8: notes 621 and some notes. I believe this will turn up sonie of the Same results Kank-you Christine Vendors and cost where applicable Type of Search NA Sequence (#) Dialog AA Sequence (#) Searcher Phone #: Questel/Orbit Structure (#) Searcher Location: Dr.Link Bibliographic Date Searcher Picked Up: Litigation Date Completed: Sequence Systems Fulltext Searcher Prep & Review Time: Patent Family Clerical Prep Time: Other Online Time:

PTO-1590 (8-01)

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    FILE 'HCAPLUS' ENTERED AT 11:57:20 ON 28 AUG 2002
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L2
           360 S INOMATA H?/AU
L3
          2835 S KOJIMA M?/AU
           514 S SUDO Y?/AU
L4
L5
           112 S SESHIMOTO O?/AU
        - 3873 S L1-5
1.6
L7 .
            70 S L6 AND FLUORESCEN?
L8 :
            6 S L7 AND NUCLEOTID?
L9
         89880 S ?CYANIN? OR ?STYRYL?
L10
             1 S L8 AND L9
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    L12
L13
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L16
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L17
           373 S L16 AND L13-15
               E SULFON/CT
               E SULFONATES+ALL/CT
         18654 S E2, E46-47
L18
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               E PHOSPHATES, BIOLOGICAL STUDIES+ALL/CT
          6154 S E3
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             2 S L22 NOT L12
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L25
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L26
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L31
             5 S L30
L32
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#### MAUPIN 09/829,467

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               STR L32
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               SAVE TEMP L36 MAU467P/A
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L38
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L53
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L55
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L57
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L62
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L64
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            8 S L65 AND L66
L67
            18 S L66 NOT L67 18 cites
L68
            7 S L67 NOT L12
L69
                               7 cites
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5Th for all 5Th sourching

VAR G1=O/S/11
REP G2=(2-9) C
NODE ATTRIBUTES:
CONNECT IS E3 RC AT 8
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

L36 45490 SEA FILE=REGISTRY SSS FUL L34

#### MAUPIN 09/829,467

#### => d ibib abs hitstr 1

L29 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1996:437966 HCAPLUS

DOCUMENT NUMBER: 125:81266

TITLE: Dye-biomolecule conjugates as contrast agents for

in-vivo near-IR diagnostic methods

INVENTOR(S): Licha, Kai; Riefke, Bjoern; Semmler, Wolfhard; Speck,

Ulrich; Hilger, Christoph-Stephan

PATENT ASSIGNEE(S): Institut fuer Diagnostikforschung Gmbh an der Freien

Universitaet Berlin, Germany

SOURCE: Ger. Offen., 18 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

£	PATENT NO.	KIND	DATE		APPLICATION N	ο.	DATE			
C	CA 2205906	AA A1	19960613 19960613		DE 1994-44450 CA 1995-22059 WO 1995-DE146	06	19951010			
					GB, GR, IE, IT,	LU	MC, NL,	PT,	SE	
P					AU 1995-37409					
P	U 709152	В2	19990819							
E	P 796111	A1	19970924		EP 1995-93534	8	19951010			
	R: AT, B	E, CH, DE	, DK, ES,	FR,	GB, GR, IE, IT,	LI,	LU, MC,	NL,	PT,	SE
C	N 1174511	Α	19980225		CN 1995-19662	4	19951010			
. Н	เบ 77378	A2	19980428		ни 1997-1797		19951010			
J	P 10510250	T2	19981006		HU 1997-1797 JP 1995-51722	8	19951010			
· J	TP 2002012782	A2	20020115		JP 2001-14390	6	19951010			
E	P 1181940	A2	20020227		JP 2001-14390 EP 2001-25036	6	19951010	•		
E	P 1181940	A3	20020313							
	R: AT, B	E, CH, DE	, DK, ES,	FR,	GB, GR, IT, LI,	LU,	, NL, SE,	MC,	PT,	ΙE
Z	A 9509707	A	19960529		ZA 1995-9707		19951115			
N	io 9702509.	A	19970602		NO 1997-2509 US 1997-84936		19970602			
Ü	IS 6083485	Α	20000704		US 1997-84936	9	19971107			
					US 2001-85066					
	ORITY APPLN. INFO.:									
					EP 1995-935348	ДΊ	19951010			
					JP 1995-517228 WO 1995-DE1465 US 1997-849369	A2	19951010			
					WO 1995-DE1465	W	19951010			
					US 1997-849369	A1	19971107			
					US 2000-518947					
SMILLE	COURCE (C).	3.47	DDM 105	0100	_					

OTHER SOURCE(S):

MARPAT 125:81266

GΙ

MeO<sub>2</sub>C Me Me Me CO<sub>2</sub>Me No So<sub>3</sub>-
$$\left\{CH_2\right\}_3$$
 CH<sub>2</sub> CH<sub>2</sub>

Ι

AB Conjugates B.scriptl.(FWn)m [B = biol. recognition mol. (mol. wt. .ltoreq.30,000); F = dye; W = hydrophilic group to improve water soly.; .scriptl. = 0-6; n = 0-10; m = 1-100] are useful as contrast agents in fluorescent and transillumination diagnostic procedures in vivo. Recognition mol. B may bind specifically to selected cell populations or receptors, may be a nonspecifically binding macromol., or may become enriched in tissues, tumors, or blood. Thus, cyanine dye I was administered i.v. to mice bearing tumor LS174T. I became enriched in the tumor after 18 h, as shown by IR fluorescence during irradn. at 740 nm.

IT 22195-47-7 51143-35-2, Glutaconaldehyde dianil

hydrochloride 80566-25-2 178698-85-6 178822-76-9 178822-77-0 178822-78-1

RL: RCT (Reactant); RACT (Reactant or reagent)

(dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

RN 22195-47-7 HCAPLUS

CN 1,3-Dioxolane-4-methanamine, 2,2-dimethyl- (9CI) (CA INDEX NAME)

RN 51143-35-2 HCAPLUS

Ph-N=CH-CH2-CH=CH-CH=N-Ph

#### ●x HCl

RN 80566-25-2 HCAPLUS

CN 1H-Benz[e]indolium, 3-ethyl-1,1,2-trimethyl-, iodide (9CI) (CA INDEX NAME)

• I.

RN 178698-85-6 HCAPLUS

CN Glycine, N-[2-[bis(carboxymethyl)amino]ethyl]-N-[2-(2,6-dioxo-4-morpholinyl)ethyl]-, monoethyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 106145-40-8 CMF C14 H21 N3 O9

CM 2

CRN 64-17-5 CMF C2 H6 O

 $_{\rm H3C}-_{\rm CH2}-_{\rm OH}$ 

RN 178822-76-9 HCAPLUS

CN 3H-Indolium, 5-(carboxymethyl)-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-1-ethyl-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

Et 
$$(CH_2)_4 - SO_3H$$

$$N^+ CH = CH - CH = CH - CH$$

$$Me$$

$$Me$$

$$Me$$

$$Me$$

$$Me$$

RN 178822-77-0 HCAPLUS

CN 3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-5-[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, sodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

RN 178822-78-1 HCAPLUS

CN 2,5-Pyrrolidinedione, 1-[[(2,3,3-trimethyl-3H-indol-5-yl)acetyl]oxy](9CI) (CA INDEX NAME)

178822-68-9P 178822-69-0P 178822-70-3P ΙT

178822-71-4P 178822-72-5P 178822-73-6P

178822-74-7P 178822-75-8P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

RN

178822-68-9 HCAPLUS
3H-Indolium, 5-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-2-[7-[5-[2-CN  $[(2,5-{\tt dioxo-l-pyrrolidinyl})\,{\tt oxy}]\,-2-{\tt oxoethyl}]\,-1,\,3-{\tt dihydro-3},\,3-{\tt dimethyl-l-(4-1)}$ sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-mathematical description of the sulfobutyl) and the sulfobutyl) are sulfobutyl) and sulfobutyl) are sulfobutyl).sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 178822-69-0 HCAPLUS

CN 3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2ylidene]-1,3,5-heptatrienyl]-1-ethyl-3,3-dimethyl-5-(2-hydrazino-2oxoethyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

$$\begin{array}{c|c} & \text{Et} \\ & \\ & \\ \text{N}^+ \\ \text{CH} \\ \text{$$

PAGE 1-B

178822-70-3 HCAPLUS RN

CN Heptanoic acid, 5-methyl-6-(phenylhydrazono)- (9CI) (CA INDEX NAME)

RN

178822-71-4 HCAPLUS
3H-Indolium, 3-(3-carboxypropyl)-2,3-dimethyl-1-(4-sulfobutyl)-, inner CN salt, sodium salt (9CI) (CA INDEX NAME)

Na

RN 178822-72-5 HCAPLUS

CN 1,2-Cyclobutanedione, 3-ethoxy-4-[(3-ethyl-1,3-dihydro-1,1-dimethyl-2Hbenz[e]indol-2-ylidene)methyl]- (9CI) (CA INDEX NAME)

178822-73-6 HCAPLUS RN

1,2-Cyclobutanedione, 3-[(3-ethyl-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-CN 2-ylidene)methyl]-4-hydroxy- (9CI) (CA INDEX NAME)

RN 178822-74-7 HCAPLUS

CN 3H-Indole-5-acetamide, N-[(2,2-dimethyl-1,3-dioxolan-4-yl)methyl]-2,3,3-trimethyl-, potassium salt (9CI) (CA INDEX NAME)

K

RN 178822-75-8 HCAPLUS

CN 3H-Indolium, 5-[2-[[(2,2-dimethyl-1,3-dioxolan-4-yl)methyl]amino]-2-oxoethyl]-2,3,3-trimethyl-1-(4-sulfobutyl)-, inner salt, potassium salt (9CI) (CA INDEX NAME)

Me O 
$$CH_2-NH-C-CH_2$$
 Me Me Me Me Me (CH2)  $4-SO_3-$ 

K

IT 178822-60-1DP, conjugates with biomols. 178822-61-2DP,
 conjugates with biomols. 178822-62-3DP, conjugates with biomols.
 178822-63-4DP, conjugates with biomols. 178822-64-5DP,
 conjugates with biomols. 178822-65-6DP, conjugates with biomols.
 178822-66-7DP, conjugates with biomols. 178822-67-8DP,
 conjugates with biomols.
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological)

study); PREP (Preparation); USES (Uses)
 (dye-biomol. conjugates as contrast agents for in-vivo near-IR
 diagnostic methods)

RN 178822-60-1 HCAPLUS

CN 3H-Indolium, 5-[2-[(1,2-dicarboxyethyl)amino]-2-oxoethyl]-2-[7-[5-[2-[(1,2-dicarboxyethyl)amino]-2-oxoethyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-

sulfobutyl)-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

PAGE 1-A

K

PAGE 1-B

RN 178822-61-2 HCAPLUS

CN 3H-Indolium, 5-[14-carboxy-7,10,13-tris(carboxymethyl)-2-oxo-3,4,7,10,13-pentaazatetradec-1-yl]-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-1-ethyl-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 178822-62-3 HCAPLUS

CN Poly(oxy-1,2-ethanediy1), .alpha.-hydro-.omega.-methoxy-, ether with 2-[7-[1,3-dihydro-5-[2-[(2-hydroxyethy1)amino]-2-oxoethoxy]-3,3-dimethy1-1-(4-sulfobuty1)-2H-indol-2-ylidene]-1,3,5-heptatrieny1]-5-[2-[(2-hydroxyethy1)amino]-2-oxoethoxy]-3,3-dimethy1-1-(4-sulfobuty1)-3H-indolium inner salt (2:1), monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

$$= CH - CH = CH - CH = CH - CH = CH - CH_{2} -$$

PAGE 1-C

$$-CH_2-CH_2$$
 OMe

RN 178822-63-4 HCAPLUS

CN Poly(oxy-1,2-ethanediy1), .alpha.-[2-[[[2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobuty1)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobuty1)-3H-indolium-5-yl]carbonyl]amino]ethyl]-.omega.-methoxy-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

$$-O3S - (CH2)4$$

$$N^{+} CH = CH$$

$$CH2 - CH2 - CH2 - OH2 - CH2 - NH - C$$

$$Me$$

$$Me$$

$$Me$$

Na

PAGE 1-B

RN 178822-64-5 HCAPLUS

CN 3H-Indolium, 3-(3-carboxypropyl)-2-[7-[3-(3-carboxypropyl)-1,3-dihydro-3-methyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3-methyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

Na

RN 178822-65-6 HCAPLUS

CN 1H-Benz[e]indolium, 1-(3-carboxypropyl)-2-[[3-[[3-(3-carboxypropyl)-1,3-dihydro-3-methyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]methyl]-4-oxo-2-cyclobuten-1-ylidene]methyl]-3-ethyl-1-methyl-, inner salt (9CI) (CA INDEX NAME)

RN 178822-66-7 HCAPLUS

CN 3H-Indolium, 5-[2-[(2,3-dihydroxypropyl)amino]-2-oxoethyl]-2-[7-[5-[2-[(2,3-dihydroxypropyl)amino]-2-oxoethyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

$$= CH \xrightarrow{N^{+}} O OH OH CH_{2}-CH-CH_{2}-OH$$

RN 178822-67-8 HCAPLUS

CN 3H-Indolium, 2-[7-[1,3-dihydro-5-(methoxycarbonyl)-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-5-(methoxycarbonyl)-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, sodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

-OMe

RN 58-85-5 HCAPLUS

CN 1H-Thieno[3,4-d]imidazole-4-pentanoic acid, hexahydro-2-oxo-, (3aS,4S,6aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

=> d ind

L29 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2002 ACS

IC ICM A61K049-00

ICS G01N021-64; C09B069-10; C09B023-02; C09B023-14

CC 9-5 (Biochemical Methods)

Section cross-reference(s): 28, 41

ST IR fluorescence dye conjugate diagnosis; cyanine dye IR fluorescence diagnosis; biomol dye conjugate fluorescence diagnosis

IT Chelating agents

(conjugates with biomols. and dyes; dye-biomol. conjugates as contrast

#### MAUPIN 09/829,467

```
agents for in-vivo near-IR diagnostic methods)
IT
    Alcohols, biological studies
    Carboxylic acids, biological studies
    Esters, biological studies
    Ethers, biological studies
       Sulfonic acids, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates with biomols. and dyes; dye-biomol. conjugates as contrast
        agents for in-vivo near-IR diagnostic methods)
IT
    Dyes
       Dyes, cyanine
        (conjugates with biomols.; dye-biomol. conjugates as contrast agents
        for in-vivo near-IR diagnostic methods)
IT
    Carotenes and Carotenoids
    Lymphokines and Cytokines
    Neurohormones
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugates with dyes; dye-biomol. conjugates as contrast agents for
        in-vivo near-IR diagnostic methods)
IT
    Diagnosis
        (dye-biomol. conjugates as contrast agents for in-vivo near-IR
        diagnostic methods)
ΙT
    Animal tissue
    Blood
    Neoplasm
        (enrichment in; dye-biomol. conjugates as contrast agents for in-vivo
        near-IR diagnostic methods)
ΙT
    Animal cell
        (specific binding to; dye-biomol. conjugates as contrast agents for
        in-vivo near-IR diagnostic methods)
ΙT
    Receptors
    RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (specific binding to; dye-biomol. conjugates as contrast agents for
        in-vivo near-IR diagnostic methods)
TT
    Enzymes
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (substrates for, conjugates with dyes; dye-biomol. conjugates as
        contrast agents for in-vivo near-IR diagnostic methods)
IT
    Spectrochemical analysis
        (IR, dye-biomol. conjugates as contrast agents for in-vivo near-IR
        diagnostic methods)
ΙT
    Ligands
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (conjugated, with dyes; dye-biomol. conjugates as contrast agents for
        in-vivo near-IR diagnostic methods)
    Agglutinins and Lectins
    Amino acids, biological studies
    Animal growth regulators
    Antigens
    Carbohydrates and Sugars, biological studies
    Coenzymes
    Haptens
    Hormones
    Macromolecular compounds
    Oligosaccharides
     Peptides, biological studies
     Polysaccharides
     Toxins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
```

#### MAUPIN 09/829,467

(conjugates, with dyes; dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

IT Neurohormones

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (neurotransmitters, conjugates, with dyes; dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

IT Nucleotides

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (oligo-, conjugates, with dyes; dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

IT 56-84-8, L-Aspartic acid, reactions 1633-83-6, 1,4-Butanesultone 5231-87-8 **22195-47-7 51143-35-2**, Glutaconaldehyde dianil hydrochloride 57998-45-5, 5-Methyl-6-oxoheptanoic acid 80506-64-5 **80566-25-2 178698-85-6 178822-76-9** 178822-77-0 178822-78-1

RL: RCT (Reactant); RACT (Reactant or reagent)
 (dye-biomol. conjugates as contrast agents for in-vivo near-IR
 diagnostic methods)

IT 178822-68-9P 178822-69-0P 178822-70-3P 178822-71-4P 178822-72-5P 178822-73-6P 178822-74-7P 178822-75-8P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(dye-biomol. conjugates as contrast agents for in-vivo near-IR
diagnostic methods)

IT 178822-60-1DP, conjugates with biomols. 178822-61-2DP,
 conjugates with biomols. 178822-62-3DP, conjugates with biomols.
 178822-63-4DP, conjugates with biomols. 178822-64-5DP,
 conjugates with biomols. 178822-65-6DP, conjugates with biomols.
 178822-66-7DP, conjugates with biomols. 178822-67-8DP,
 conjugates with biomols.
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

IT 58-85-5D, Biotin, conjugates with dyes 9004-54-0D, Dextran, conjugates with dyes 25322-68-3D, conjugates with biomols. and dyes RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (dye-biomol. conjugates as contrast agents for in-vivo near-IR diagnostic methods)

#### => d ibib abs hitstr 2

L29 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1996:194739 HCAPLUS

DOCUMENT NUMBER: 124:225822

TITLE: N-heteroaromatic ion and iminium ion substituted

cyanine dyes for use as fluorescence labels

INVENTOR(S): Lee, Linda G.; Woo, Sam L. PATENT ASSIGNEE(S): Biometric Imaging, Inc., USA

SOURCE: PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

	PATENT NO.			KIND DATE					A	PPLI	CATI	٥.	DATE					
	WO 9600902			A1 19960111				WO 1995-US8778						19950629				
		W:	AM,	AT,	ΑU,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CZ,	DE,	DK,	EE,	ES,	FI,
			GB,	GE,	HU,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	ΚZ,	LK,	LR,	LT,	LU,	LV,	MD,
			MG,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	ТJ,
			TT,	UA														
		RW:	ΚE,	MW,	SD,	SZ,	UG,	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,
			LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	ML,	MR,	NE,
			SN,	TD,	ΤG													
	US 5453505			A 19950926			US 1994-268852 1994063											
		9530					1996	-							1995			
	EΡ	7691	45		Α	1	1997	0423		E	P 19	95-9	2627	2	1995	0629		
		R:	AT,	BE,	CH,	DE,	ES,	FR,	GB,	IE,	ΙΤ,	LI,	LU,	ΝL,	SE			
PRIORITY APPLN. INFO.:							US 1994-268852 19940630											
	U								US 1995-388607									
						WO 1995-US8778 19950629												

OTHER SOURCE(S): MARPAT 124:225822
GI For diagram(s), see printed CA Issue.

AΒ The present invention relates to iminium ion-substituted cyanine dyes having a fluorescence absorbance of between about 500 and 900 nm, a reduced tendency to aggregate and enhanced photostability. The cyanine dyes of the present invention are represented by formula I where n is 0, 1, 2 or 3; R1 and R2 are taken together to form an arom. ring or a fused polycyclic arom. ring; R3 and R4 are taken together to form an arom. ring or a fused polycyclic arom. ring; R5 and R6 are independently selected from the group consisting of (CH2)pX where p is 1-18 and X is a functional group that reacts with amino, hydroxy and sulfhydryl nucleophiles; R7 and R8 are independently selected from the group consisting of H, C1-C10 alkyl groups and where R7 and R8 are taken together to form a 5- or 6-membered heterocyclic ring; R9 are each independently selected from the group consisting of H, alkyl and where >1 R9 are taken together to form a 5- or 6-membered ring; Y is selected from the group consisting of C(CH3)2, S, O and Se; and Z is selected from the group consisting of C(CH3)2, S, O and Se. The present invention also relates to a method for using the cyanine dyes of the present invention to fluorescent label mols., particularly biomols. such as antibodies, DNA, carbohydrates and cells.

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IT 174829-15-3P 174829-16-4P 174829-17-5P 174829-18-6P 174829-20-0P 174829-21-1P 174829-23-3P 174829-25-5P 174829-27-7P 174829-28-8P 174829-29-9P 174829-30-2P 174829-32-4P 174829-33-5P 174829-34-6P 174829-35-7P 174829-37-9P 174829-39-1P
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174829-40-4P 174829-41-5P 174829-42-6P 174829-44-8P 174829-45-9P 174829-46-0P 174829-47-1P 174829-49-3P 174829-51-7P 174829-52-8P 174829-53-9P 174829-54-0P 174829-56-2P 174829-57-3P 174829-58-4P 174829-59-5P 174829-61-9P

RL: ARG (Analytical reagent use); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

RN 174829-15-3 HCAPLUS

CN

3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-16-4 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-(3-methyl-1H-imidazolium-1-yl)-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-17-5 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(1-pyrrolidinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-18-6 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(4-morpholinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-20-0 HCAPLUS

CN 1H-Benz[e]indolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-21-1 HCAPLUS

CN 1H-Benz[e]indolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]ethylidene]-2-(3-methyl-1H-imidazolium-1-yl)-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-23-3 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclopenten-1-yl]ethenyl]-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-25-5 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-6-fluoro-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-6-fluoro-3,3-dimethyl-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-27-7 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-28-8 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-(3-methyl-1H-imidazol-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} \text{Me} \\ \text{(CH}_2)_5 - \text{CO}_2 - \\ \text{N} \\ \text{CH} - \text{CH} - \text{CH} - \text{CH} - \text{CH} \\ \end{array}$$

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*
RN 174829-29-9 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-[4-(1-pyrrolidinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

$$(CH_2)_5 - CO_2 - CH_2)_5$$
 $CH = CH - CH - CH$ 

RN 174829-30-2 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-[4-(4-morpholinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

$$(CH_2)_5 - CO_2 - CH_2)_5$$
 $CH = CH - CH_2$ 

RN 174829-32-4 HCAPLUS

CN Naphtho[2,1-d]thiazolium, 3-(5-carboxypentyl)-2-[2-[3-[3-(5-carboxypentyl)naphtho[2,1-d]thiazol-2(3H)-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-33-5 HCAPLUS

CN Naphtho[2,1-d]thiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphtho[2,1-d]thiazol-2(3H)-ylidene]ethylidene]-2-(3-methyl-1H-imidazol-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-34-6 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-chloro-1-cyclopenten-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-35-7 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclopenten-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-37-9 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzothiazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-39-1 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-40-4 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-(3-methyl-1H-imidazol-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-41-5 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-[4-(1-pyrrolidinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-42-6 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-[4-(4-morpholinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-44-8 HCAPLUS

CN Naphtho[2,1-d]selenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphtho[2,1-d]selenazol-2(3H)-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-45-9 HCAPLUS

CN Naphtho[2,1-d]selenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphtho[2,1-d]selenazol-2(3H)-ylidene]ethylidene]-2-(3-methyl-1H-imidazol-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-46-0 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-chloro-1-cyclopenten-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-47-1 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclopenten-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-49-3 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzoselenazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-51-7 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-52-8 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-(3-methyl-1H-imidazolium-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-53-9 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-[4-(1-pyrrolidinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

$$(CH_2)_5 - CO_2 - V_+ - O_2C - (CH_2)_5$$
 $CH = CH - CH - CH$ 

RN 174829-54-0 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-([3-(5-carboxypentyl)-2(3H)-

benzoxazolylidene]ethylidene]-2-[4-(4-morpholinyl)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

$$(CH_2)_5 - CO_2 - O_2C - (CH_2)_5$$
 $CH = CH - CH - CH$ 

RN 174829-56-2 HCAPLUS

CN Naphth[2,1-d]oxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphth[2,1-d]oxazol-2(3H)-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-57-3 HCAPLUS

CN Naphth[2,1-d]oxazolium, 3-(5-carboxypenty1)-2-[2-[3-[[3-(5-carboxypenty1)naphth[2,1-d]oxazol-2(3H)-ylidene]ethylidene]-2-(3-methyl-1H-imidazolium-1-yl)-1-cyclohexen-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

\*\*\* FRAGMENT DIAGRAM IS INCOMPLETE \*\*\*

RN 174829-58-4 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-chloro-1-cyclopenten-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-59-5 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclopenten-1-yl]ethenyl]-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 174829-61-9 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzoxazolylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, bis(inner salt) (9CI) (CA INDEX NAME)

109-97-7DP, Pyrrole, cyanine dyes contg. 110-86-1DP, ΙT Pyridine, cyanine dyes contg. 120-73-0DP, Purine, cyanine dyes contg. 151-56-4DP, Aziridine, cyanine dyes contg. 288-13-1DP, Pyrazole, cyanine dyes contg. 289-80-5DP, Pyridazine, cyanine dyes contg. 289-95-2DP, Pyrimidine, cyanine dyes contg. 290-37-9DP, Pyrazine, cyanine dyes contg. 541-59-3DP, 1H-Pyrrole-2,5-dione, cyanine dyes contg. 1122-58-3DP, cyanine dyes contg. 2767-91-1DP, 4-(4-Morpholinyl)pyridine, cyanine dyes contg. 2831-66-5DP, cyanine dyes contg. 6153-86-2DP, cyanine dyes contg. 16969-45-2DP, Pyridinium, cyanine dyes contg. 17009-89-1DP , 1-Methylimidazolium, cyanine dyes contg. 17009-90-4DP, Imidazolium, cyanine dyes contg. 22559-70-2DP, Quinolinium, cyanine dyes contg. 23715-85-7DP, Isoquinolinium, cyanine dyes contg. 82436-78-0DP, cyanine dyes contg. 104302-69-4DP 3-(2-Pyridyldithio)propionamide, cyanine dyes contg. RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses) (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells) 109-97-7 HCAPLUS RN CN 1H-Pyrrole (9CI) (CA INDEX NAME)



RN 110-86-1 HCAPLUS CN Pyridine (6CI, 7CI, 8CI, 9CI) (CA INDEX NAME)



RN 120-73-0 HCAPLUS CN 1H-Purine (9CI) (CA INDEX NAME)

RN 151-56-4 HCAPLUS CN Aziridine (9CI) (CA INDEX NAME)

H N \_\_\_\_

RN 288-13-1 HCAPLUS CN 1H-Pyrazole (9CI) (CA INDEX NAME)



RN 289-80-5 HCAPLUS CN Pyridazine (8CI, 9CI) (CA INDEX NAME)



RN 289-95-2 HCAPLUS CN Pyrimidine (8CI, 9CI) (CA INDEX NAME)



RN 290-37-9 HCAPLUS CN Pyrazine (8CI, 9CI) (CA INDEX NAME)



RN 541-59-3 HCAPLUS CN 1H-Pyrrole-2,5-dione (9CI) (CA INDEX NAME)

RN 1122-58-3 HCAPLUS

CN 4-Pyridinamine, N, N-dimethyl- (9CI) (CA INDEX NAME)

RN 2767-91-1 HCAPLUS

CN Morpholine, 4-(4-pyridinyl)- (9CI) (CA INDEX NAME)

RN 2831-66-5 HCAPLUS

CN 1,3,5-Triazine, 2,4-dichloro- (9CI) (CA INDEX NAME)

RN 6153-86-2 HCAPLUS

CN - 1,3,5-Triazine, 2-chloro- (9CI) (CA INDEX NAME)

RN 16969-45-2 HCAPLUS

CN Pyridine, conjugate acid (8CI, 9CI) (CA INDEX NAME)



H+

RN 17009-89-1 HCAPLUS CN 1H-Imidazole, 1-methyl-, conjugate monoacid (9CI) (CA INDEX NAME)



H+

RN 17009-90-4 HCAPLUS CN 1H-Imidazole, conjugate monoacid (9CI) (CA INDEX NAME)



H+

RN 22559-70-2 HCAPLUS CN Quinoline, conjugate acid (8CI, 9CI) (CA INDEX NAME)



● H-

RN 23715-85-7 HCAPLUS CN Isoquinoline, conjugate acid (8CI, 9CI) (CA INDEX NAME)

H+

RN 82436-78-0 HCAPLUS

CN 3-Pyrrolidinesulfonic acid, 1-hydroxy-2,5-dioxo- (9CI) (CA INDEX NAME)

RN 104302-69-4 HCAPLUS

CN Propanamide, 3-(2-pyridinyldithio)- (9CI) (CA INDEX NAME)

IT **54849-69-3**, IR 144 **144377-05-9**, Cy5

RL: PRP (Properties)

(N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

RN 54849-69-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-1,1-dimethyl-3-(3-sulfopropyl)-2H-benz[e]indol-2-ylidene]ethylidene]-2-[4-(ethoxycarbonyl)-1-piperazinyl]-1-cyclopenten-1-yl]ethenyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 55660-40-7

CMF C50 H58 N4 O8 S2

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 144377-05-9 HCAPLUS

CN 3H-Indolium, 5-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-2-[5-[5-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, sodium salt (9CI) (CA INDEX NAME)

# PAGE 1-A

Na

PAGE 1-B

62-53-3, Benzenamine, reactions 616-47-7, N-Methylimidazole 1122-58-3 1640-39-7 6066-82-6, N-Hydroxysuccinimide 41532-84-7 RL: RCT (Reactant); RACT (Reactant or reagent) (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

62-53-3 HCAPLUS RN

Benzenamine (9CI) (CA INDEX NAME) CN

616-47-7 HCAPLUS RN 1H-Imidazole, 1-methyl- (9CI) (CA INDEX NAME) CN

RN1122-58-3 HCAPLUS CN 4-Pyridinamine, N, N-dimethyl- (9CI) (CA INDEX NAME)



RN 1640-39-7 HCAPLUS 3H-Indole, 2,3,3-trimethyl- (6CI, 7CI, 8CI, 9CI) (CA INDEX NAME) CN

RN 6066-82-6 HCAPLUS CN 2,5-Pyrrolidinedione, 1-hydroxy- (9CI) (CA INDEX NAME)

RN 41532-84-7 HCAPLUS CN 1H-Benz[e]indole, 1,1,2-trimethyl- (9CI) (CA INDEX NAME)

63857-00-1P 171429-39-3P 174829-14-2P IT 174829-19-7P 174829-22-2P 174829-24-4P 174829-26-6P 174829-31-3P 174829-36-8P 174829-38-0P 174829-43-7P 174829-48-2P 174829-50-6P 174829-55-1P 174829-60-8P 174829-62-0P 174829-63-1P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells) 63857-00-1 HCAPLUS RN Benzenamine, N-[[2-chloro-3-[(phenylamino)methylene]-1-cyclohexen-1-CN yl]methylene]-, monohydrochloride (9CI) (CA INDEX NAME)

HC1

RN 171429-39-3 HCAPLUS

CN 1H-Benz[e]indolium, 3-(5-carboxypentyl)-1,1,2-trimethyl-, bromide (9CI) (CA INDEX NAME)

Me Me Me 
$$+N$$
HO<sub>2</sub>C- (CH<sub>2</sub>) 5

● Br-

RN 174829-14-2 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 174829-19-7 HCAPLUS

CN 1H-Benz[e]indolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 174829-22-2 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-chloro-1-cyclopenten-1-yl]ethenyl]-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 174829-24-4 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[2-[3-[[1-(5-carboxypentyl)-6-fluoro-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-6-fluoro-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 174829-26-6 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzothiazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-31-3 HCAPLUS

CN Naphtho[2,1-d]thiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphtho[2,1-d]thiazol-2(3H)-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-36-8 HCAPLUS

CN Benzothiazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzothiazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, inner salt (9CI) (CA INDEX NAME)

RN 174829-38-0 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoselenazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-,
inner salt (9CI) (CA INDEX NAME)

RN 174829-43-7 HCAPLUS

CN Naphtho[2,1-d]selenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphtho[2,1-d]selenazol-2(3H)-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-48-2 HCAPLUS

CN Benzoselenazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzoselenazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, inner salt (9CI) (CA INDEX NAME)

RN 174829-50-6 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-2(3H)-benzoxazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl}-, inner salt (9CI) (CA INDEX NAME)

RN 174829-55-1 HCAPLUS

CN Naphth[2,1-d]oxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)naphth[2,1-d]oxazol-2(3H)-ylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-, inner salt (9CI) (CA INDEX NAME)

RN 174829-60-8 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[2-[3-[[3-(5-carboxypentyl)-5-fluoro-2(3H)-benzoxazolylidene]ethylidene]-2-chloro-1-cyclohexen-1-yl]ethenyl]-5-fluoro-, inner salt (9CI) (CA INDEX NAME)

RN 174829-62-0 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2,3,3-trimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 174829-63-1 HCAPLUS

CN 3H-Indolium, 2-[2-[3-[[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]ethylidene]-2-[4-(dimethylamino)pyridinio]-1-cyclohexen-1-yl]ethenyl]-1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3,3-dimethyl-, mono(inner salt) (9CI) (CA INDEX NAME)

=> d ind 2

L29 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2002 ACS

IC ICM G01N033-58

ICS C09B023-02; C09B023-00

CC 9-5 (Biochemical Methods)

Section cross-reference(s): 28

ST cyanine dye fluorescent label biopolymer cell; heteroarom ion substituted cyanine dye probe; iminium ion substituted cyanine dye label

IT Cell

## Dyes, cyanine

Photolysis

Ultraviolet and visible spectra

(N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

IT Antibodies

Biopolymers

Carbohydrates and Sugars, reactions

Deoxyribonucleic acids

RL: RCT (Reactant); RACT (Reactant or reagent)

(N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

IT Functional groups

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (azidonitrophenyl, cyanine dyes contg.; N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as fluorescent labels for biopolymers and cells)

IT Acid halides

Aldehydes, preparation

Azides

Carboxylic acids, preparation

Cyanates

Disulfides

```
Esters, preparation
      Sulfonic acids, preparation
     RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (cyanine dyes contg.; N-heteroarom. ion- and iminium ion-substituted
        cyanine dyes prepn. as fluorescent labels for biopolymers and cells)
IT
     Iminium compounds
    RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (ions, cyanine dyes substituted with; N-heteroarom. ion- and iminium
        ion-substituted cyanine dyes prepn. as fluorescent labels for
        biopolymers and cells)
IT
     Immunoglobulins
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (G, N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn.
        as fluorescent labels for biopolymers and cells)
    Heterocyclic compounds
    RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (arom., ions, cyanine dyes substituted with; N-heteroarom. ion- and
        iminium ion-substituted cyanine dyes prepn. as fluorescent labels for
       biopolymers and cells)
IT
    Molecules
        (biochem., N-heteroarom. ion- and iminium ion-substituted cyanine dyes
        prepn. as fluorescent labels for biopolymers and cells)
    Esters, preparation
IT
    RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (imido, cyanine dyes contg.; N-heteroarom. ion- and iminium
        ion-substituted cyanine dyes prepn. as fluorescent labels for
        biopolymers and cells)
IT
    Fluorescent substances
        (probes, N-heteroarom. ion- and iminium ion-substituted cyanine dyes
        prepn. as fluorescent labels for biopolymers and cells)
    174829-15-3P 174829-16-4P 174829-17-5P
TT
    174829-18-6P 174829-20-0P 174829-21-1P
    174829-23-3P 174829-25-5P 174829-27-7P
    174829-28-8P 174829-29-9P 174829-30-2P
    174829-32-4P 174829-33-5P 174829-34-6P
    174829-35-7P 174829-37-9P 174829-39-1P
    174829-40-4P 174829-41-5P 174829-42-6P
    174829-44-8P 174829-45-9P 174829-46-0P
    174829-47-1P 174829-49-3P 174829-51-7P
    174829-52-8P 174829-53-9P 174829-54-0P
    174829-56-2P 174829-57-3P 174829-58-4P
    174829-59-5P 174829-61-9P
    RL: ARG (Analytical reagent use); PRP (Properties); RCT (Reactant); SPN
     (Synthetic preparation); ANST (Analytical study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as
        fluorescent labels for biopolymers and cells)
ΙT
     107-22-2DP, Glyoxal, cyanine dyes contg. 109-97-7DP, Pyrrole,
     cyanine dyes contq. 110-86-1DP, Pyridine, cyanine dyes contq.
    120-73-0DP, Purine, cyanine dyes contg. 151-56-4DP,
    Aziridine, cyanine dyes contg. 288-13-1DP, Pyrazole, cyanine
    dyes contg. 289-80-5DP, Pyridazine, cyanine dyes contg.
    289-95-2DP, Pyrimidine, cyanine dyes contg. 290-37-9DP,
     Pyrazine, cyanine dyes contg. 302-01-2DP, Hydrazine, cyanine dyes contg.
        541-59-3DP, 1H-Pyrrole-2,5-dione, cyanine dyes contg.
     1122-58-3DP, cyanine dyes contg. 2767-91-1DP,
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### MAUPIN 09/829,467

```
4-(4-Morpholinyl)pyridine, cyanine dyes contg. 2831-66-5DP,
     cyanine dyes contg. 6153-86-2DP, cyanine dyes contg.
     7704-34-9DP, Sulfur, cyanine dyes contg.
                                                7782-44-7DP, Oxygen, cyanine
     dyes contq.
                   7782-49-2DP, Selenium, cyanine dyes contg.
    16969-45-2DP, Pyridinium, cyanine dyes contg. 17009-89-1DP
     , 1-Methylimidazolium, cyanine dyes contg. 17009-90-4DP,
     Imidazolium, cyanine dyes contg. 22559-70-2DP, Quinolinium,
     cyanine dyes contg. 23715-85-7DP, Isoquinolinium, cyanine dyes
    contg. 82436-78-0DP, cyanine dyes contg. 104302-69-4DP
     , 3-(2-Pyridyldithio)propionamide, cyanine dyes contg.
    RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as
        fluorescent labels for biopolymers and cells)
     54849-69-3, IR 144 144377-05-9, Cy5
                                           169799-14-8, Cy7
IT
     RL: PRP (Properties)
        (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as
        fluorescent labels for biopolymers and cells)
     62-53-3, Benzenamine, reactions
                                      68-12-2, DMF, reactions
     108-94-1, Cyclohexanone, reactions 616-47-7, N-Methylimidazole
    1122-58-3 1640-39-7
                           4224-70-8, 6-Bromohexanoic acid
     6066-82-6, N-Hydroxysuccinimide
                                      10025-87-3, Phosphorus
    oxychloride 41532-84-7
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as
        fluorescent labels for biopolymers and cells)
ΙT
     63857-00-1P 171429-39-3P 174829-14-2P
    174829-19-7P 174829-22-2P 174829-24-4P
    174829-26-6P 174829-31-3P 174829-36-8P
    174829-38-0P 174829-43-7P 174829-48-2P
    174829-50-6P 174829-55-1P 174829-60-8P
    174829-62-0P 174829-63-1P
    RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (N-heteroarom. ion- and iminium ion-substituted cyanine dyes prepn. as
        fluorescent labels for biopolymers and cells)
ΙT
    302-04-5P, Isothiocyanate, preparation
    RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
     (Analytical study); PREP (Preparation); USES (Uses)
        (cyanine dyes contg.; N-heteroarom. ion- and iminium ion-substituted
        cyanine dyes prepn. as fluorescent labels for biopolymers and cells)
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Inventor Sourch

#### MAUPIN 09/829,467

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L12 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2002 ACS 2001:753071 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

135:303873

TITLE:

Fluorescent labeled nucleotides,

synthesis and application as probes and primers

INVENTOR(S):

Shinoki, Hiroshi; Inomata, Hiroko; Kojima, Masayoshi; Sudo, Yukio;

Seshimoto, Osamu

PATENT ASSIGNEE(S):

Fuji Photo Film Co., Ltd., Japan Jpn. Kokai Tokkyo Koho, 14 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2001288197	A2	20011016	JP 2000-107675	20000410
US 2002064782	A1	20020530	US 2001-829467	20010409
EP 1152008	A2	20011107	EP 2001-107864	20010410
EP 1152008	A3	20020320		

152008 A3 20020320 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

PRIORITY APPLN. INFO.:

JP 2000-107675 A 20000410

MARPAT 135:303873 OTHER SOURCE(S): AΒ The present invention provides a fluorescent substance which is represented by a formula: A-B-C wherein A is a residue of natural or synthetic nucleotide, oligonucleotide, polynucleotide, or deriv. thereof, and binds to B at a base moiety in said residue; B is a divalent linking group or a single bond; and C is a deriv. of fluorescent dye having 0 or 1 sulfonate or phosphate moiety. Fluorescent dye is cyanine, melocyanine, or styryl. Preferably A is AMP, ADP, ATP, GMP, GDP, GTP, CMP, CDP, CTP, UMP, UDP, UTP, TMP, TDP, TTP, 2-Me-AMP, 2-Me-ADP, 2-Me-ATP, 1-Me-GMP, 1-Me-GDP, 1-Me-GTP, 5-Me-CMP, 5-Me-CDP, 5-Me-CTP, 5-MeO-CMP, 5-MeO-CDP, 5-MeO-CTP. B is preferably -CH2-, -CH=CH-, triple bond, -CO-, -O-, -S-, -NH-, or aminoaryl. Synthesis of labeled nucleic acids using the nucleotides via reverse transcription, terminal transferase

reaction, random prime method, PCR, or nick translation, is claimed. fluorescent substance of the present invention is useful as label for nucleic acids, reagent for detecting nucleic acids, or diagnostic reagent. Kits for nucleic acid detection are claimed. Synthesis of 8 indolenine cyanine compds. and conjugation with dUTP, and use for DNA probe prepn., are described.

ΙT 23065-05-6, Styryl

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (fluorescent labeled nucleotide synthesis and application as probes and primers)

RN 23065-05-6 HCAPLUS

CN Ethenyl, 2-phenyl- (9CI) (CA INDEX NAME)

HC== CH- Ph

IT 366451-16-3P 366451-17-4P 366451-18-5P 366451-19-6P 366451-20-9P 366451-21-0P

#### 366451-22-1P 366451-23-2P

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); SPN (Synthetic preparation); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)

(fluorescent labeled nucleotide synthesis and

application as probes and primers)

RN

366451-16-3 HCAPLUS
3H-Indolium, 1-(5-carboxypentyl)-2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-2H-CN indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl- (9CI) (CA INDEX NAME)

RN 366451-17-4 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-5-chloro-2-[5-[5-chloro-1,3-dihydro-1-[2-(2-hydroxyethoxy)ethyl]-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl- (9CI) (CA INDEX NAME)

RN 366451-18-5 HCAPLUS

3H-Indolium, 5-(aminosulfonyl)-2-[3-[5-(aminosulfonyl)-1-(5-carboxypentyl)-CN 1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3dimethyl- (9CI) (CA INDEX NAME)

366451-19-6 HCAPLUS RN

3H-Indolium, 1-(5-carboxypentyl)-5-chloro-2-[3-[5-chloro-1,3-dihydro-1-[2-CN (2-hydroxyethoxy)ethyl]-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-3,3dimethyl- (9CI) (CA INDEX NAME)

HO<sub>2</sub>C- (CH<sub>2</sub>) 5 
$$CH_2$$
- CH<sub>2</sub>- CH<sub>2</sub>- OH  $CH_2$ - OH  $CH_2$ - CH<sub>2</sub>- OH  $CH_2$ - OH  $CH_$ 

RN

366451-20-9 HCAPLUS 3H-Indolium, 5-(aminosulfonyl)-2-[5-[5-(aminosulfonyl)-1-[6-[[2-[1-[2-CN deoxy-5-0-[hydroxy([hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5pyrimidinyl]ethenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-, inner salt, trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

Na

PAGE 1-B

366451-21-0 HCAPLUS RN

3H-Indolium, 5-chloro-2-[5-[5-chloro-1-[6-[[2-[1-[2-deoxy-5-0-CN

[hydroxy[[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]ethenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-[2-(2-hydroxyethoxy)ethyl]-3,3-dimethyl-, inner salt, trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-A

●3 Na

PAGE 1-B

RN 366451-22-1 HCAPLUS

CN 3H-Indolium, 5-(aminosulfonyl)-2-[3-[5-(aminosulfonyl)-1-[6-[[2-[1-[2-deoxy-5-0-[hydroxy[[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]ethenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-, inner salt, trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

●3 Na

### PAGE 1-B

RN 366451-23-2 HCAPLUS

CN 3H-Indolium, 5-chloro-2-[3-[5-chloro-1-[6-[[2-[1-[2-deoxy-5-O-' [hydroxy[[hydroxy(phosphonooxy)phosphinyl]]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]ethenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1-[2-(2-hydroxyethoxy)ethyl]-3,3-dimethyl-, inner salt, trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

●3 Na

PAGE 1-B

56-65-5, 5'-ATP, reactions 58-64-0, 5'-ADP, reactions ΙT 58-97-9, 5'-UMP, reactions 58-98-0, 5'-UDP, reactions 61-19-8, 5'-AMP, reactions 63-37-6, CMP 63-38-7 , CDP 63-39-8, 5'-UTP 65-47-4, 5'-CTP 85-32-5 , 5'-GMP **86-01-1**, 5'-GTP **95-50-1**, 1, 2-Dichloro benzene 122-51-0, Triethyl orthoformate 146-91-8, 5'-GDP 365-07-1, DTMP 365-08-2, TTP 491-97-4 , TDP 628-89-7 1173-82-6, DUTP 1173-82-6D, DUTP, aminoaryl 1927-31-7, DATP 2056-98-6, DCTP 2564-35-4, DGTP 3590-36-1 4224-70-8, 6-Bromo hexanoic acid 14315-97-0 20309-92-6 25981-83-3 39923-67-6 39923-68-7, 2-Methyl-ADP 42467-24-3 , 2-Methyl-ATP 52940-67-7 52988-98-4 76528-21-7 80677-38-9 112242-04-3 130536-69-5 327174-86-7 366451-24-3 RL: RCT (Reactant); RACT (Reactant or reagent) (fluorescent labeled nucleotide synthesis and application as probes and primers) RN 56-65-5 HCAPLUS Adenosine 5'-(tetrahydrogen triphosphate) (8CI, 9CI) (CA INDEX NAME) CN

$$NH2$$
 $NH2$ 
 $NH2$ 

RN 58-64-0 HCAPLUS

CN Adenosine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 58-97-9 HCAPLUS

CN 5'-Uridylic acid (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 58-98-0 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

RN 61-19-8 HCAPLUS

CN 5'-Adenylic acid (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 63-37-6 HCAPLUS

CN 5'-Cytidylic acid (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 63-38-7 HCAPLUS

CN Cytidine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 63-39-8 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate) (8CI, 9CI) (CA INDEX NAME)

RN 65-47-4 HCAPLUS

CN Cytidine 5'-(tetrahydrogen triphosphate) (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 85-32-5 HCAPLUS

CN 5'-Guanylic acid (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 86-01-1 HCAPLUS

CN Guanosine 5'-(tetrahydrogen triphosphate) (8CI, 9CI) (CA INDEX NAME)

RN 95-50-1 HCAPLUS

CN Benzene, 1,2-dichloro- (9CI) (CA INDEX NAME)

RN 122-51-0 HCAPLUS

CN Ethane, 1,1',1''-[methylidynetris(oxy)]tris- (9CI) (CA INDEX NAME)

RN 146-91-8 HCAPLUS

CN Guanosine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 365-07-1 HCAPLUS

CN 5'-Thymidylic acid (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 365-08-2 HCAPLUS

CN Thymidine 5'-(tetrahydrogen triphosphate) (8CI, 9CI) (CA INDEX NAME)

RN 491-97-4 HCAPLUS

CN Thymidine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 628-89-7 HCAPLUS

CN Ethanol, 2-(2-chloroethoxy) - (6CI, 7CI, 8CI, 9CI) (CA INDEX NAME)

 ${\tt C1CH_2-CH_2-O-CH_2-CH_2-OH}$ 

RN 1173-82-6 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 1173-82-6 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy- (9CI) (CA INDEX NAME)

RN 1927-31-7 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 2'-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 2056-98-6 HCAPLUS

CN Cytidine 5'-(tetrahydrogen triphosphate), 2'-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 2564-35-4 HCAPLUS

CN Guanosine 5'-(tetrahydrogen triphosphate), 2'-deoxy- (9CI) (CA INDEX NAME)

RN 3590-36-1 HCAPLUS CN 5'-Cytidylic acid, 5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 4224-70-8 HCAPLUS

CN Hexanoic acid, 6-bromo- (6CI, 7CI, 8CI, 9CI) (CA INDEX NAME)

 $Br-(CH_2)_5-CO_2H$ 

RN 14315-97-0 HCAPLUS

CN Propane, 1,1,3-trimethoxy- (9CI) (CA INDEX NAME)

RN 20309-92-6 HCAPLUS

CN 5'-Guanylic acid, 1-methyl- (9CI) (CA INDEX NAME)

Me N N N 
$$R$$
 R R OPO3H2

RN 25981-83-3 HCAPLUS CN 3H-Indole, 5-chloro-2,3,3-trimethyl- (8CI, 9CI) (CA INDEX NAME)

RN 39923-67-6 HCAPLUS CN 5'-Adenylic acid, 2-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 39923-68-7 HCAPLUS CN Adenosine 5'-(trihydrogen diphosphate), 2-methyl- (9CI) (CA INDEX NAME)

RN 42467-24-3 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 2-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 52940-67-7 HCAPLUS

CN Guanosine 5'-(tetrahydrogen triphosphate), 1-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Me N N N O OH OPO3H2
$$R R S$$
 OH OPO3H2
 $R S O OH$ 

RN 52988-98-4 HCAPLUS

CN Guanosine 5'-(trihydrogen diphosphate), 1-methyl- (9CI) (CA INDEX NAME)

RN 76528-21-7 HCAPLUS

CN 5'-Cytidylic acid, 5-methoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 80677-38-9 HCAPLUS

CN Cytidine 5'-(trihydrogen diphosphate), 5-methoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 112242-04-3 HCAPLUS

CN Cytidine 5'-(trihydrogen diphosphate), 5-methyl- (9CI) (CA INDEX NAME)

RN 130536-69-5 HCAPLUS

CN Ethanol, 2-(2-iodoethoxy)- (9CI) (CA INDEX NAME)

ICH2-CH2-O-CH2-CH2-OH

RN 327174-86-7 HCAPLUS

CN Cytidine 5'-(tetrahydrogen triphosphate), 5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 366451-24-3 HCAPLUS

CN Cytidine 5'-(tetrahydrogen triphosphate), 5-methoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 366451-26-5DP, bromide 366451-27-6DP, bromide
366451-28-7DP, bromide

### MAUPIN 09/829,467

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(fluorescent labeled nucleotide synthesis and

application as probes and primers)

RN

366451-26-5 HCAPLUS
3H-Indolium, 5-(aminosulfonyl)-1-(5-carboxypentyl)-2,3,3-trimethyl- (9CI) CN (CA INDEX NAME)

RN 366451-27-6 HCAPLUS

3H-Indolium, 1-(5-carboxypentyl)-5-chloro-2,3,3-trimethyl- (9CI) (CA CN INDEX NAME)

C1 Me Me Me 
$$_{\rm N^+}$$
 (CH<sub>2</sub>) 5-CO<sub>2</sub>H

366451-28-7 HCAPLUS RN

3H-Indolium, 5-chloro-1-[2-(2-hydroxyethoxy)ethyl]-2,3,3-trimethyl- (9CI) CN (CA INDEX NAME)

ΙT **75-03-6**, Ethyl iodide **62306-05-2** 

RL: RCT (Reactant); RACT (Reactant or reagent) (reactant; fluorescent labeled nucleotide synthesis and application as probes and primers)

RN 75-03-6 HCAPLUS

Ethane, iodo- (8CI, 9CI) (CA INDEX NAME) CN

 ${
m H}_3{
m C}-{
m CH}_2-{
m I}$ 

RN62306-05-2 HCAPLUS

3H-Indole-5-sulfonamide, 2,3,3-trimethyl- (9CI) (CA INDEX NAME) CN

$$\begin{array}{c|c} & & & & \\ & & & \\ H_2N-S & & & \\ \parallel & & & \\ O & & & \\ \end{array}$$

IT 366451-25-4DP, iodide

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(reactant; fluorescent labeled nucleotide synthesis

and application as probes and primers)

RN 366451-25-4 HCAPLUS

CN 3H-Indolium, 5-(aminosulfonyl)-1-ethyl-2,3,3-trimethyl- (9CI) (CA INDEX NAME)

IC ICM C07H019-10

ICS C07H019-20; C07H021-00; C09K011-06; C12N015-09; C12Q001-68; G01N033-58; C07D209-08; C07D209-30; C07D403-06; C07D403-14

CC 28-1 (Heterocyclic Compounds (More Than One Hetero Atom))

Section cross-reference(s): 3, 9

ST fluorescent labeled nucleotide synthesis probe primer;

cyanine melocyanine styryl nucleotide

synthesis probe primer

IT Diagnosis

(agents; fluorescent labeled nucleotide synthesis

and application as probes and primers)

IT Phosphates, biological studies

Sulfonates

RL: BOC (Biological occurrence); BSU (Biological study, unclassified);

BIOL (Biological study); OCCU (Occurrence)

(dye contg.; fluorescent labeled nucleotide

synthesis and application as probes and primers)

IT Cyanine dyes

Fluorescent dyes

Test kits

(fluorescent labeled nucleotide synthesis and

application as probes and primers)

IT Nucleotides, preparation

Oligonucleotides

Polynucleotides

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); SPN (Synthetic preparation); ANST (Analytical study); BIOL (Biological study);

PREP (Preparation); USES (Uses)

(fluorescent labeled nucleotide synthesis and

application as probes and primers)

IT Nucleic acid amplification (method)

### MAUPIN 09/829,467

6,

```
(terminal transferase reaction, use in labeled nucleic acid synthesis;
        fluorescent labeled nucleotide synthesis and
        application as probes and primers)
IT
     PCR (polymerase chain reaction)
     Reverse transcription
        (use in labeled nucleic acid synthesis; fluorescent labeled
        nucleotide synthesis and application as probes and primers)
ΙT
     23065-05-6, Styryl
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (fluorescent labeled nucleotide synthesis and
        application as probes and primers)
     366451-16-3P 366451-17-4P 366451-18-5P
     366451-19-6P 366451-20-9P 366451-21-0P
     366451-22-1P 366451-23-2P
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); SPN
     (Synthetic preparation); ANST (Analytical study); BIOL (Biological study);
     PREP (Preparation); USES (Uses)
        (fluorescent labeled nucleotide synthesis and
        application as probes and primers)
     56-65-5, 5'-ATP, reactions 58-64-0, 5'-ADP, reactions 58-97-9, 5'-UMP, reactions 58-98-0, 5'-UDP, reactions
     61-19-8, 5'-AMP, reactions 63-37-6, CMP 63-38-7
     CDP 63-39-8, 5'-UTP 65-47-4, 5'-CTP 85-32-5, 5'-GMP 86-01-1, 5'-GTP 95-50-1, 1, 2-Dichloro benzene 122-51-0, Triethyl orthoformate 146-91-8,
     5'-GDP 365-07-1, DTMP 365-08-2, TTP 491-97-4
       TDP 628-89-7 1173-82-6, DUTP 1173-82-6D,
     DUTP, aminoaryl 1927-31-7, DATP 2056-98-6, DCTP
     2564-35-4, DGTP 3590-36-1 4224-70-8, 6-Bromo
     hexanoic acid 14315-97-0 20309-92-6 25981-83-3
     39923-67-6 39923-68-7, 2-Methyl-ADP 42467-24-3
       2-Methyl-ATP 52940-67-7 52988-98-4
     76528-21-7 80677-38-9 112242-04-3
     130536-69-5 327174-86-7 366451-24-3
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (fluorescent labeled nucleotide synthesis and
        application as probes and primers)
TT
     366451-26-5DP, bromide 366451-27-6DP, bromide
     366451-28-7DP, bromide
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (fluorescent labeled nucleotide synthesis and
        application as probes and primers)
IΤ
     75-03-6, Ethyl iodide 62306-05-2
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (reactant; fluorescent labeled nucleotide synthesis
        and application as probes and primers)
     366451-25-4DP, iodide
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (reactant; fluorescent labeled nucleotide synthesis
        and application as probes and primers)
```

Species search for p. 21 of spec which corresponds to structure on MAUPIN 09/829,467 => d his (FILE 'HOME' ENTERED AT 14:35:22 ON 28 AUG 2002) FILE 'REGISTRY' ENTERED AT 14:35:35 ON 28 AUG 2002 ACT MAU467P/A STR

p. 2-6 og the inventor search. L145490 SEA FILE=REGISTRY SSS FUL L1 parint search L2 STR L1 L3 23 S L3 SSS SAM SUB=L2 L4541 S L3 SSS FUL SUB=L2 L5

SAVE L5 MAU467S1/A 43 S L5 AND (NCNC2-NCNC3/ES OR NCNC3/ES) = 5 ub set search L6

FILE 'HCAPLUS' ENTERED AT 14:44:42 ON 28 AUG 2002 L7 30 S L6 3 S (1996:437966 OR 1996:194739 OR 2001:753071)/AN - in ventos work  $\Gamma8$ L9 29 S L7 NOT L8 10 S L9 AND PATENT/DT L10 getting rid of bod dates 9 S L10 AND PRD<20010409/ L11

19 S L9 NOT PATENT/DT L12 L13 17 S L12 AND PD<20010409

26 S L11 OR L13 **26** L14 S 366451-20-9/REG#

FILE 'REGISTRY' ENTERED AT 14:55:09 ON 28 AUG 2002 L15 1 S 366451-20-9/RN

FILE 'HCAPLUS' ENTERED AT 14:55:10 ON 28 AUG 2002 1 S L15 S 366451-18-5/REG#

FILE 'REGISTRY' ENTERED AT 14:55:56 ON 28 AUG 2002 1 S 366451-18-5/RN L17

FILE 'HCAPLUS' ENTERED AT 14:55:56 ON 28 AUG 2002 1 S L17

L18 1 S L16 OR L18 T.19

L16

FILE 'REGISTRY' ENTERED AT 14:58:07 ON 28 AUG 2002 L20 498 S L5 NOT L6

L21 4 S L20 AND N=2 AND CL=2 AND O=4 2 S L21 AND "HYDROXYETHOXY" L22

3 S L20 AND N=4 AND S=2 AND O=6 L23

1 S L23 AND C31 H41 N4 O6 S2/MF

L25 1 S L20 AND C=33 AND N=4 AND O=6 AND S=2

FILE 'HCAPLUS' ENTERED AT 15:06:02 ON 28 AUG 2002 applicant's work L26 1 S L22 OR L24

applicant's
sperific cpds
- they are
found only in inventors citation

```
=> d que 120
L1 STR ANY CYPLO

2 7 13
C. 3 G1 8 G2 Cy Me C Me
1 C C 14 10 @11 12

6 C. C 4 9
5

VAR G1=0/S/11
```

REP G2=(2-9) C
NODE ATTRIBUTES:
CONNECT IS E3 RC AT 8
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

## GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 14

### STEREO ATTRIBUTES: NONE

L2 45490 SEA FILE=REGISTRY SSS FUL L1 L3 STR

Me 15

16 Me

2

17, C, 19, C, 23

2

10, C, 3, C, 8, Ak

10, C, 7, C, N, 25

Ak~C=0

29, 30, 31

& species STR.

(C t+z) n NODE ATTRIBUTES:

CONNECT IS E3 RC AT 8
DEFAULT MLEVEL IS ATOM
GGCAT IS LIN UNS AT 13
GGCAT IS LIN SAT AT 29
DEFAULT ECLEVEL IS LIMITED

# GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 24

### STEREO ATTRIBUTES: NONE

L5 541 SEA FILE=REGISTRY SUB=L2 SSS FUL L3

L6 43 SEA FILE=REGISTRY ABB=ON PLU=ON L5 AND (NCNC2-NCNC3/ES OR

NCNC3/ES)

L20 498 SEA FILE=REGISTRY ABB=ON PLU=ON L5 NOT L6

=> d ibib abs hitstr

L14 ANSWER 1 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2002:449855 HCAPLUS

DOCUMENT NUMBER:

137:30254

TITLE:

Fluorescent labeling of protein C-terminal with puromycin analogs linked to fluorophores and high-throughput assay technologies for in vitro

analysis of protein interactions

INVENTOR(S):

Yanagawa, Hiroshi; Doi, Nobuhide; Miyamoto, Etsuko;

Takashima, Hideaki; Oyama, Rieko

PATENT ASSIGNEE(S):

Keio University, Japan PCT Int. Appl., 95 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_\_ \_\_\_\_ -----\_\_\_\_\_ WO 2002046395 A1 20020613 WO 2001-JP10731 20011207 <--

W: CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

PT, SE, TR PRIORITY APPLN. INFO.: JP 2000-373105 A 20001207 <--

A method for modifying protein C-terminal with a reagent which contains an acceptor region having a group capable of binding to a protein through a transpeptidation reaction and a modifying region contg. a modifier linked to the acceptor region via a nucleotide linker, is disclosed. A template contg. an ORF encoding a protein, a 5'-unntranslated region (UTR) contg. a promoter and an enhancer located in the 5'-side of the ORF and a 3'-terminal region contg. a PolyA sequence located in the 3'-side of the ORF is expressed to thereby synthesize a protein. The protein thus synthesized is then purified. The yield of the modified protein in the protein C-terminal modification method can be largely improved and protein interactions can be detected at an improved level in the method of detecting interactions among various mols. The authors developed and tested a simple method for fluorescence labeling and interaction anal. of proteins based on a highly efficient in vitro translation system combined with high-throughput technologies such as microarrays and fluorescence cross-correlation spectroscopy (FCCS). By use of puromycin analogs linked to various fluorophores through a deoxycytidylic acid linker, a single fluorophore can be efficiently incorporated into a protein at the carboxyl terminus during in vitro translation. The authors confirmed that the resulting fluorescently labeled proteins are useful for probing protein-protein and protein-DNA interactions by means of pulldown assay, DNA microarrays, and FCCS in model expts. These fluorescence assay systems can be easily extended to highly parallel anal. of protein interactions in studies of functional genomics. Interactions involving c-Fos, c-Jun, and DNA were studied by labeling with rhodamine green or Cy5 using puromycin-contg. modifying agents.

ΙT 436083-90-8 436083-91-9

> RL: MOA (Modifier or additive use); RGT (Reagent); RACT (Reactant or reagent); USES (Uses)

(fluorescence labeling of protein C-terminal with puromycin analogs linked to fluorophores and high-throughput assay technol. for in vitro anal. of protein interactions)

RN 436083-90-8 HCAPLUS CN Adenosine, 2'-deoxy-5-[3-[[6-[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]hexyl]amino]-3-oxo-1-propenyl]-5'-O-[21-[(3aS,4S,6aR)hexahydro-2-oxo-1H-thieno[3,4-d]imidazol-4-yl]-1-hydroxy-1-oxido-10,17-dioxo-2-oxa-9,16-diaza-1-phosphaheneicos-1-yl]uridylyl-(3'.fwdarw.5')-2'-deoxycytidylyl-(3'.fwdarw.5')-3'-[[(2S)-2-amino-3-(4-methoxyphenyl)-1-oxopropyl]amino]-3'-deoxy-N,N-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-B

PAGE 1-C

RN 436083-91-9 HCAPLUS

CN Adenosine, 5'-O-[[[6-[[5-[(3aS,4S,6aR)-2-amino-3a,4,6,6a-tetrahydro-1H-thieno[3,4-d]imidazol-4-yl]-1-oxopentyl]amino]hexyl]oxy]hydroxyphosphinyl]-2'-deoxy-5-[3-[[6-[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]hexyl]amino]-3-oxo-1-propenyl]uridylyl-(3'.fwdarw.5')-2'-deoxycytidylyl-(3'.fwdarw.5')-3'-[[(2S)-2-amino-3-(4-methoxyphenyl)-1-oxopropyl]amino]-3'-deoxy-N,N-dimethyl-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A MeO

PAGE 1-B

REFERENCE COUNT:

12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 2

L14 ANSWER 2 OF 26 HCAPLUS COPYRIGHT 2002 ACS 2002:405773 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

136:403155

TITLE:

Monofunctional indocyanine labeling reagents and

improved method for their production

Caputo, Giuseppe; Della, Ciana Leopoldo INVENTOR(S):

PATENT ASSIGNEE(S):

Innosense S.R.L., Italy Eur. Pat. Appl., 25 pp.

SOURCE: CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

KIND DATE PATENT NO. APPLICATION NO. DATE ---------\_\_\_\_\_\_ EP 2000-126019 20001128 EP 1209205 A1 20020529 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR EP 1211294 A1 20020605 EP 2001-127884 20011123 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR US 2002065421 Α1 20020530 US 2001-995350 20011127 <--AU 2001-93440 AU 2001093440 Α5 20020606 20011127 <--PRIORITY APPLN. INFO.: EP 2000-126019 A 20001128 <--OTHER SOURCE(S): CASREACT 136:403155; MARPAT 136:403155 A process for prepg. an asym. indocyanine dye is characterized in that it AΒ comprises the steps of (a) reacting a first quaternized indolenine or substituted indolenine with RN:CH(CH:CX)nNHR or hydrochloride thereof (n = solvent selected from the group consisting of acetic acid, acetic

0, 1 R = Ph or substituted Ph, X H, halogen or alkyl, preferably Cl) in a anhydride and mixts. thereof in the presence of acetyl chloride, to obtain an intermediate hemicyanine; and (b) further reacting this intermediate hemicyanine with a second quaternized indolenine or substituted indolenine different from the first indolenine. The process is characterized by high yields, readily obtained starting materials, and facile workup. The products are suitable as fluorescent labels emitting in the IR and near-IR region which can be excited by means of simple light-emitting or laser diodes and have high extinction coeffs. Thus, a hemicyanine was prepd. from 1-ethyl-2,3,3-trimethylindolenium iodide, malonaldehyde dianil, and acetyl chloride; this product was then treated with 1-(3-acetoxypropyl)-2,3,3-trimethylindolenium iodide and then with 2-cyanoethyl tetraisopropylphosphorodiamidite to provide a fluorescent labeling dye.

431943-84-9P 431943-86-1P IT

> RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); PREP (Preparation); USES (Uses)

(dye; prodn. of monofunctional indocyanine fluorescent labeling dyes)

RN 431943-84-9 HCAPLUS

1H-Benz[e]indolium, 3-[6-[[3-[1-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonoox CN y)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4tetrahydro-2,4-dioxo-5-pyrimidinyl]-2-propenyl]amino]-6-oxohexyl]-2-[3-(1ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1-propenyl]-1,1dimethyl-6,8-disulfo-, inner salt, hexasodium salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 431943-86-1 HCAPLUS

CN lH-Benz[e]indolium, 3-[6-[[3-[1-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonoox y)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-2-propenyl]amino]-6-oxohexyl]-2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-, inner salt, pentasodium salt (9CI) (CA INDEX NAME)

HO3S

PAGE 1-B

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 3

L14 ANSWER 3 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:730822 HCAPLUS

135:283541

TITLE:

A novel polypeptide-protein 11 of growth

hormone-family and a polynucleotide sequence encoding

the same

INVENTOR(S):

Mao, Yumin; Xie, Yi

PATENT ASSIGNEE(S):

Shanghai Biowindow Gene Development Inc., Peop. Rep.

APPLICATION NO.

DATE

China

SOURCE:

PCT Int. Appl., 35 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

KIND DATE

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.

	WO 2001072832			A1 20011004			WO 2001-CN489						20010326 <							
		W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CO,		
			CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,		
			HU,	ID,	ΙL,	IN,	IS,	JP,	KΕ,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,		
			LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NO,	NZ,	PL,	PT,	RO,	RU,		
			SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	ΤZ,	UA,	UG,	US,	UZ,	VN,		
			YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	MT						
		RW:	GH,	GM,	KE,	LS,	MW,	MΖ,	SD,	SL,	SZ,	ΤZ,	UG,	ZW,	AT,	BE,	CH,	CY,		
			DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙΤ,	LU,	MC,	NL,	PT,	SE,	TR,	BF,		
			ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG				
CN 1315391 A 20011003 CN 2000-115180 20000327																				
PRIORITY APPLN. INFO.: CN 2000-115180 A 20000327 <																				
AB	AB The present invention discloses a novel polypeptide-protein 11 of growth																			
	hormone-family and a polynucleotide encoding the same, as well as a method																			
	of producing the polypeptide by DNA recombinant technique. The present																			
	invention also discloses methods of using the polypeptide in treatment of																			
	various diseases, such as malignant tumor, blood disease, HIV infection,														•					
	immunol. disease, various inflammations and so on. The present invention																			
																		eutic		
													such	nov	el p	olynı	ıcle	otide		
		codin					_					-								
ΙT	158613-48-0, Cy3-dUTP 158613-49-1, Cy5-dUTP																			

RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(label; protein 11 of growth hormone-family, analogs, antagonists, promoters, inhibitors, encoding polynucleotides, and antibodies for diagnosis and treatment of cancer, blood disease, HIV, immunol. disease and inflammation)

RN 158613-48-0 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-CN[(2E)-3-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2-propenylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]- (9CI) (CA INDEX NAME)

PAGE 1-B

RN 158613-49-1 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E,4E)-5-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2,4-pentadienylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]-(9CI) (CA INDEX NAME)

HO

PAGE 1-B

4

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

# => d ibib abs hitstr 4

L14 ANSWER 4 OF 26 HCAPLUS COPYRIGHT 2002 ACS 2001:713386 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

135:271898

TITLE:

A novel polypeptide-human CDC4 analogous protein and

the polynucleotide encoding said polypeptide and

antagonistic antibody

INVENTOR(S):

Mao, Yumin; Xie, Yi

PATENT ASSIGNEE(S):

Biowindow Gene Development Inc., Peop. Rep. China

SOURCE:

PCT Int. Appl., 38 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
PATENT NO.
                          KIND DATE
                                                       APPLICATION NO. DATE
      WO 2001070779 A1 20010927 WO 2001-CN157 20010226 <--
            W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CR,
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BI, BZ, CA, CH, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                             A 20010912
      CN 1312284
                                                        CN 2000-111935 20000307
PRIORITY APPLN. INFO.:
                                                     CN 2000-111935 A 20000307 <--
AB
      The invention discloses a new kind of polypeptide-human CDC4 analogous
      protein 12 and the polynucleotide encoding said polypeptide and a process
      for producing the polypeptide by recombinant methods. It also discloses
      the method of applying the polypeptide for the treatment of various kinds
      of diseases, such as cancer, hemopathy, HIV infection, immune diseases and
      inflammation. The antagonist of the polypeptide and therapeutic use of
      the same is also disclosed. In addn., it refers to the use of
      polynucleotide encoding said human CDC4 analogous protein 12.
ΙT
      158613-48-0 158613-49-1
      RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
       (Analytical study); BIOL (Biological study); USES (Uses)
           (label; human CDC4 analogous protein, encoding polynucleotide,
          antibody, and antagonist for diagnostic and therapeutic uses)
      158613-48-0 HCAPLUS
RN
      Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-
CN
       [(2E)-3-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2-propenylidene]-2,3-dihydro-
```

Absolute stereochemistry. Double bond geometry as shown.

(CA INDEX NAME)

3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]- (9CI)

PAGE 1-B

RN 158613-49-1 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E,4E)-5-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2,4-pentadienylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]-(9CI) (CA INDEX NAME)

HO

PAGE 1-B

3

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 4

L14 ANSWER 4 OF 26 HCAPLUS COPYRIGHT 2002 ACS 2001:713386 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

135:271898

TITLE:

A novel polypeptide-human CDC4 analogous protein and

the polynucleotide encoding said polypeptide and

antagonistic antibody

INVENTOR(S):

Mao, Yumin; Xie, Yi

PATENT ASSIGNEE(S):

Biowindow Gene Development Inc., Peop. Rep. China

SOURCE:

PCT Int. Appl., 38 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

DATE														
20010226 <														
CR,														
HU,														
LU,														
SD,														
YU,														
CY,														
BF,														
ous														
protein 12 and the polynucleotide encoding said polypeptide and a process for producing the polypeptide by recombinant methods. It also discloses														
the method of applying the polypeptide for the treatment of various kinds														
of diseases, such as cancer, hemopathy, HIV infection, immune diseases and														
inflammation. The antagonist of the polypeptide and therapeutic use of														
the same is also disclosed. In addn., it refers to the use of polynucleotide encoding said human CDC4 analogous protein 12.														
Γ														
-deoxy-5-[(1E)-3-[[6-[(2E)-2-														
ihydro-														
(9CI)														

Absolute stereochemistry. Double bond geometry as shown.

(CA INDEX NAME)

PAGE 1-B

RN 158613-49-1 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E,4E)-5-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2,4-pentadienylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]-(9CI) (CA INDEX NAME)

HO3S 
$$\frac{CH_2)_5}{Me}$$
  $\frac{E}{Me}$   $\frac{E}{Me}$ 

HO

PAGE 1-B

3

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

### => d ibib abs hitstr 5

L14 ANSWER 5 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:575379 HCAPLUS

DOCUMENT NUMBER:

135:341009

TITLE:

Rapid functional analysis of protein-protein

interactions by fluorescent C-terminal labeling and

single-molecule imaging

AUTHOR(S):

Yamaguchi, J.; Nemoto, N.; Sasaki, T.; Tokumasu, A.;

Mimori-Kiyosue, Y.; Yagi, T.; Funatsu, T.

CORPORATE SOURCE:

Department of Physics, School of Science and

Engineering, Waseda University, Shinjuku-ku, Tokyo,

169-8555, Japan

SOURCE:

FEBS Letters (2001), 502(3), 79-83 CODEN: FEBLAL; ISSN: 0014-5793

PUBLISHER:

Elsevier Science B.V. Journal

DOCUMENT TYPE: LANGUAGE: English

Detection of protein-protein interactions is a fundamental step to AΒ understanding gene function. Here we report a sensitive and rapid method for assaying protein-protein interactions at the single-mol. level. Protein mols. were synthesized in a cell-free translation system in the presence of Cy5-puro, a fluorescent puromycin, using mRNA without a stop codon. The interaction of proteins thus prepd. was visualized using a single-mol. imaging technique. As a demonstration of this method, a motor protein, kinesin, was labeled with Cy5-puro at an efficiency of about 90%, and the processive movement of kinesin along microtubules was obsd. by using total internal reflection microscopy. It took only 2 h from the synthesis of proteins to the functional anal. This method is applicable to the functional anal. of various kinds of proteins.

ΙT 370884-42-7P

> RL: ARU (Analytical role, unclassified); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation)

(protein-protein interactions by fluorescent C-terminal labeling and single-mol. imaging)

RN 370884-42-7 HCAPLUS

CN Adenosine, 2'-deoxy-5'-O-[[2-[2-[6-(1-ethyl-1,3-dihydro-3,3dimethyl-5-sulfo-2H-indol-2-ylidene)-2,4-hexadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethoxy]ethoxy]hydroxyphosphinyl]cytidylyl-(3'.fwdarw.5')-3'-[(2S)-2-amino-3-(4-methoxyphenyl)-1-oxopropyl]amino]-3'deoxy-N, N-dimethyl-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

REFERENCE COUNT:

23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 6

L14 ANSWER 6 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:507799 HCAPLUS

DOCUMENT NUMBER: 135:93921

TITLE: Mobility-modifying cyanine dyes

INVENTOR(S): Menchen, Steven M.; Benson, Scott C.; Rosenblum,

Barnett B.; Khan, Shaheer H.

PATENT ASSIGNEE(S): PE Corporation, USA SOURCE: PCT Int. Appl., 133 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIN					ND	DATE			A	PPLI	CATI	и ис	٥.	DATE				
					_	2001			WO 2001-US152 20010103									
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	ВG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	
		ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NO,	NZ,	PL,	PT,	RO,	RU,	
		SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	UZ,	VN,	YU,	
		ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM						
	RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,	
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙΤ,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	
		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	G₩,	ML,	MR,	NE,	SN,	TD,	TG			
DRITY	ITY APPLN. INFO.:							1	US 2	000-	4772	70	Α	2000	0104	<		

PRIORITY APPLN. INFO.: US 2000-477270 A 20000104 OTHER SOURCE(S): MARPAT 135:93921

The present invention provides a novel class of fluorescent cyanine dye compds. that are modified at one of the heterocyclic ring nitrogen atoms with a mobility-modifying moiety that permits the electrophoretic mobilities of polynucleotides labeled with the mobility-modifying cyanine dyes to be adjusted or tuned in a predictable fashion while retaining enzymic activity. The ability to predictably tune the relative electrophoretic mobilities of the dyes permits the creation of sets of mobility-matched fluorescent dyes of a variety of structures for a variety of applications, including fluorescence-based 4-color nucleic acid sequencing reactions.

# IT 349491-76-5P 349491-78-7P

RL: ARG (Analytical reagent use); IMF (Industrial manufacture); TEM (Technical or engineered material use); ANST (Analytical study); PREP (Preparation); USES (Uses)

(mobility-modifying fluorescent cyanine dyes for nucleic acid sequencing reactions)

RN 349491-76-5 HCAPLUS

CN lH-Benz[e]indolium, 2-[5-[3-[6-[[3-[4-amino-1,2-dihydro-2-oxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-[(2,4-disulfophenyl)methyl]-1,1-dimethyl-, inner salt (9CI) (CA INDEX NAME)

Me Me Me 
$$(CH_2)_5$$
  $N_H$   $C = C$ 

PAGE 1-B

PAGE 2-A

SO3H

RN 349491-78-7 HCAPLUS

CN

1H-Benz[e]indolium, 2-[5-[1,3-dihydro-1,1-dimethyl-3-[6-oxo-6-[[2-[[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]oxy]ethyl]amino]hexyl]-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt

# (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

Me Me Me 
$$(CH_2)_5$$
  $N_H$   $(CH_2)_3$   $SO_3$ 

# PAGE 1-B

# MAUPIN 09/829,467

=> d ibib abs hitstr 7

L14 ANSWER 7 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:295103 HCAPLUS

DOCUMENT NUMBER: 136:1249

TITLE: Detection of mitochondrial single nucleotide

polymorphisms using a primer elongation reaction on

oligonucleotide microarrays

AUTHOR(S): Erdogan, Fikret; Kirchner, Roland; Mann, Wolfgang;

Ropers, Hans-Hilger; Nuber, Ulrike A.

CORPORATE SOURCE: Max-Planck Institute for Molecular Genetics, Berlin,

14195, Germany

SOURCE: Nucleic Acids Research (2001), 29(7),

e36/1-e36/7

CODEN: NARHAD; ISSN: 0305-1048

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal LANGUAGE: English

The authors have developed a novel allele-specific primer elongation AB protocol using a DNA polymerase on oligonucleotide chips. Oligonucleotide primers carrying polymorphic sites at their free 3'end were covalently bound to glass slides. The generation of single-stranded targets of genomic DNA contg. single nucleotide polymorphisms (SNPs) to be typed was achieved by an asym. PCR reaction or exonuclease treatment of phosphothicate (PTO)-modified PCR products. In the presence of DNA polymerase and all four dNTPs, with Cy3-dUTP replacing dTTP, allele-specific extension of the immobilized primers took place along a stretch of target DNA sequence. The yield of elongated products was increased by repeated reaction cycles. The authors performed multiplexed assays with many small DNA targets, or used single targets of up, to 4.4 kb mitochondrial DNA (mtDNA) sequence to detect multiple SNPs in one reaction. The latter approach greatly simplifies preamplification of SNP-contg. regions, thereby providing a framework for typing hundreds of mtDNA polymorphisms.

IT 158613-48-0, Cy3-dUTP

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(detection of mitochondrial single nucleotide polymorphisms using allele-specific elongation of immobilized oligonucleotide primers)

RN 158613-48-0 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E)-3-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2-propenylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]- (9CI) (CA INDEX NAME)

PAGE 1-B

30

REFERENCE COUNT:

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### MAUPIN 09/829,467

=> d ibib abs hitstr 8

L14 ANSWER 8 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:182053 HCAPLUS

DOCUMENT NUMBER: 135:339932

TITLE: Progress towards single-molecule sequencing: enzymatic

synthesis of nucleotide-specifically labeled DNA

AUTHOR(S): Augustin, M. A.; Ankenbauer, W.; Angerer, B.

CORPORATE SOURCE: Institut fur Biophysik und Physikalische Biochemie,

Universitat Regensburg, Regensburg, D-93051, Germany

SOURCE: Journal of Biotechnology (2001), 86(3),

289-301

CODEN: JBITD4; ISSN: 0168-1656

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

The enzymic incorporation of modified dNTPs into a growing DNA strand has intensively been studied. Modifications were detectable reporter groups such as digoxigenin or biotin, fluorochromes or aliph. side chains covalently attached to the base. Incorporation efficiencies were detd. with several DNA polymerases using linear primer-extension reactions followed by denaturing PAGE as a high-resoln. detection system. The authors describe the enzymic synthesis of DNA consisting of modified nucleotides exclusively. A defined template-primer system allows us to trace incorporation: (1) in up to 18 neighboring positions for several dUTP-derivs.; or (2) in stretches of DNA of up to 40 bases in length with complete substitution of all four natural dNTPs by differently modified counterparts. Synthesized DNA mols. are shown to particularly exhibit dramatically altered physico-chem. properties by contrast with native DNA. These results provide a fundamental data set for probe generation in single-mol. DNA sequencing (SMS).

IT 306274-02-2 306274-03-3 371920-27-3

RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses) (enzymic synthesis of nucleotide-specifically labeled DNA in relation to single-mol. sequencing)

RN 306274-02-2 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1-oxohexyl]amino]-1-propenyl]-, inner salt (9CI) (CA INDEX NAME)

-03S

PAGE 1-B

RN 306274-03-3 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[[6-[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1-oxohexyl]amino]-1-propenyl]-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

RN

371920-27-3 HCAPLUS
3H-Indolium, 1-[6-[[3-[1-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonooxy)phosp CN hinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-2-propenyl]amino]-6-oxohexyl]-2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

12

REFERENCE COUNT:

THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

#### => d ibib abs hitstr 9

AUTHOR(S):

L14 ANSWER 9 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:135278 HCAPLUS

DOCUMENT NUMBER: 135:207645

TITLE: Two-color fluorescence labeling of early and

mid-to-late replicating chromatin in living cells
Schermelleh, Lothar; Solovei, Irina; Zink, Daniele;

Cremer, Thomas

CORPORATE SOURCE: Ludwig Maximilians University, Munich, Germany

SOURCE: Chromosome Research (2001), 9(1), 77-80

CODEN: CRRSEE; ISSN: 0967-3849

PUBLISHER: Kluwer Academic Publishers

DOCUMENT TYPE: Journal LANGUAGE: English

The authors report a modified scratch-loading protocol called scratch replication labeling (SRL) for labeling chromatin in adherently growing cells. The protocol involves making scratches with a hypodermic needle tip through cells grown to subconfluency on glass coverslips. At the time of scratching, the cells are pulse-labeled in culture medium contg. Cy3-dUTP and Cy5-dUTP or Cy5-dUTP and FITC-dUTP. More efficient labeling was obtained by synchronizing the cells at the G1/S transition. Labeled cells continued to undergo mitosis and sequential labeling allowed visualization of early and mid-to-late replicating chromatin. The SRL protocol was used to label SH-EP N14 neuroblastoma cells, HeLa cells, BHK cells and primary human fibroblasts.

IT 158613-48-0, Cy3-dUTP 158613-49-1, Cy5-dUTP
RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST

(Analytical study); BIOL (Biological study); USES (Uses) (two-color fluorescence labeling of early and mid-to-late replicating chromatin in living cells)

RN 158613-48-0 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E)-3-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2-propenylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

PAGE 1-A

PAGE 1-B

RN 158613-49-1 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E,4E)-5-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2,4-pentadienylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

HO

PAGE 1-B

18

REFERENCE COUNT:

THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 10

L14 ANSWER 10 OF 26 HCAPLUS COPYRIGHT 2002 ACS

2001:38157 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 134:302397

TITLE: A comparison of optical geometries for combined flash

photolysis and total internal reflection fluorescence

microscopy

AUTHOR(S): Conibear, P. B.; Bagshaw, C. R.

Department of Biochemistry, University of Leicester, CORPORATE SOURCE:

Leicester, LE1 7RH, UK Journal of Microscopy (Oxford) (2000), SOURCE:

200(3), 218-229

CODEN: JMICAR; ISSN: 0022-2720

Blackwell Science Ltd. PUBLISHER:

DOCUMENT TYPE: Journal LANGUAGE: English

Total internal reflection fluorescence (TIRF) microscopy, used in conjunction with flash photolysis, provides a useful way of studying the kinetics of macromol. interactions. The authors compare different TIRF optical geometries to establish an optimal combination. Excitation light was introduced via 4 different arrangements: (1) a prism positioned on the microscope optical axis, (2) an offset prism with propagation through a SiO2 slide trans to the objective lens. (3) An offset prism with propagation through a SiO2 coverslip cis to a H2O-immersion objective lens and (4) a prismless arrangement using a high NA oil-immersion objective lens. Photolysis was achieved using a Xe flash lamp and a customized SiO2 condenser lens. Single myosin mols. labeled with a Cy3 fluorophore were used as a test sample. Although the offset trans prism gave the best signal-to-background ratio, a customized thin rhombic prism incorporated, on axis, into the flash condenser assembly was almost as good and was more practical for scanning multiple fields. An oil-immersion lens gave the brightest image for sample depths < 30 .mu.m but above this limit, a H2O-immersion lens was better. The prismless arrangement may offer advantages in other situations but it is important to check the actual numerical aperture of the objective lens.

ΙT 213904-25-7 213904-27-9

> RL: MOA (Modifier or additive use); PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process); USES (Uses)

(optical geometries comparison for combined flash photolysis and total internal reflection fluorescence microscopy of myosin labeled with)

RN 213904-25-7 HCAPLUS

CN dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2Hindol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

PAGE 2-A

RN 213904-27-9 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 3'-[[2-[[6-[2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-

indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

\_so3-

REFERENCE COUNT:

24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

### => d ibib abs hitstr 11

L14 ANSWER 11 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:825711 HCAPLUS

DOCUMENT NUMBER: 134:159683

TITLE: Identification of single fluorescently labelled

mononucleotide molecules in solution by spectrally

resolved time-correlated single-photon counting

AUTHOR(S): Herten, D. P.; Tinnefeld, P.; Sauer, M.

CORPORATE SOURCE: Physikalisch-Chemisches Institut, Universitat

Heidelberg, Heidelberg, 69120, Germany

SOURCE:

Applied Physics B: Lasers and Optics (2000), 71(5), 765-771

CODEN: APBOEM; ISSN: 0946-2171

PUBLISHER: Springer-Verlag

DOCUMENT TYPE: Journal LANGUAGE: English

We describe a method to identify single dye-labeled mononucleotide mols. in soln. with high classification probability based on confocal microscopy in combination with spectrally and time-resolved fluorescence detection with two detectors. For efficient excitation of the labeled mononucleotide mols. JA133-dUTP, JA169-dUTP, Cy5-dCTP, and JA242-dUTP a short-pulse diode laser emitting at 634 nm with a repetition rate of 64 MHz was applied. The time-resolved fluorescence signals of individual mols. were analyzed and identified by a max. likelihood estimator (MLE). Scatter plots of spectrally and time-resolved fluorescence data demonstrated the existence of four distinct populations with sym. shape. The distributions of each of the mononucleotide conjugates were detd. by fitting a superposition of two independent Gaussians. Taking only those single-mol. bursts which contain more than 50 photon counts, three labeled mononucleotide mols. were identified in soln. by spectrally and time-resolved fluorescence spectroscopy with a probability of correct classification of .apprxeq. 99%.

#### 325747-77-1 TT

RL: ANT (Analyte); BSU (Biological study, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study)

(identification of single fluorescently labeled mononucleotide mols. in soln. by spectrally resolved time-correlated single-photon counting)

325747-77-1 HCAPLUS RN

3H-Indolium, 2-[5-[1-[6-[[3-[4-amino-1-[2-deoxy-5-0-CN [hydroxy[[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythropentofuranosyl]-1,2-dihydro-2-oxo-5-pyrimidinyl]-2-propynyl]amino]-6oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

H2N\_

PAGE 1-B

REFERENCE COUNT:

THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 12

L14 ANSWER 12 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:814658 HCAPLUS

DOCUMENT NUMBER: 133:345552

TITLE: High-density labeling of DNA with modified or

> chromophore-tagged nucleotides using DNA polymerases Muehlegger, Klaus; Angerer, Bernhard; Seela, Frank; Ankenbauer, Waltraud; Augustin, Martin; Gumbiowski,

Karin; Zulauf, Matthias

PATENT ASSIGNEE(S): Roche Diagnostics G.m.b.H., Germany

PCT Int. Appl., 56 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

INVENTOR(S):

PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_ \_\_\_\_\_ WO 2000068422 Α2 20001116 WO 2000-EP4036 20000505 <--WO 2000068422 А3 20020404 W: JP, US RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE 20020529 EP 2000-936714 20000505 <--EP 1208230 A2 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY

PRIORITY APPLN. INFO.: EP 1999-108601 A 19990507 <--WO 2000-EP4036 W 20000505 <--AB

Subjects of the inventions are methods for enzymic DNA labeling. Nucleotides modified to carry functional or detectable groups are incorporated into newly synthesized DNA by DNA polymerases. DNA is synthesized from modified nucleoside triphosphates by DNA polymerases such that the newly synthesized DNA consists exclusively of modified nucleotides or contains modified nucleotides in high d. There are provided modified nucleoside triphosphates which are incorporated by DNA polymerases and a group of DNA polymerases which incorporate these nucleoside triphosphates in high d. Thus, modified nucleoside triphosphates, such as 7-aminopentinyl-7-deazaadenosine-2'deoxyribonucleoside-5'-triphosphate, were synthesized. Incorporation of this and other modified nucleoside triphosphates into DNA in the presence of template, primer, and Carboxydothermus hydrogenoformans, Pyrococcus, Thermococcus gorgonarius (Tgo), Pyrococcus woesei (Pwo), or a blend of Tgo and Pwo polymerases was analyzed. The combination of Tgo and Pwo polymerases seemed to be most effective.

306274-02-2P 306274-03-3P 306274-04-4P

RL: BPR (Biological process); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process)

(high-d. labeling of DNA with modified or chromophore-tagged nucleotides using DNA polymerases)

RN 306274-02-2 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[6-[2-[5-(1-CN ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1oxohexyl]amino]-1-propenyl]-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

-03S

PAGE 1-B

RN 306274-03-3 HCAPLUS

Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1-oxohexyl]amino]-1-propenyl]-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

RN 306274-04-4 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[38-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-5,12,19,26,33-pentaoxo-4,11,18,25,32-pentaazaoctatriacont-1-en-1-yl]-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

## PAGE 1-B

$$\begin{array}{c|c} \text{(CH2)5} & \\ N \\ H \end{array} \begin{array}{c} \text{(CH2)5} \\ \text{O} \end{array} \begin{array}{c} \text{(CH2)5} \\ N \\ H \end{array} \begin{array}{c} \text{(CH2)5} \\ N \\ \text{Me} \end{array} \end{array}$$

# PAGE 1-C

### => d ibib abs hitstr 13

AUTHOR(S):

L14 ANSWER 13 OF 26 HCAPLUS COPYRIGHT 2002 ACS

2000:397304 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 133:173509

Comparative single-molecule and ensemble myosin TITLE:

enzymology: sulfoindocyanine ATP and ADP derivatives Oiwa, Kazuhiro; Eccleston, John F.; Anson, Michael; Kikumoto, Mahito; Davis, Colin T.; Reid, Gordon P.; Ferenczi, Michael A.; Corrie, John E. T.; Yamada,

Akira; Nakayama, Haruto; Trentham, David R.

Communications Research Laboratory, Kansai Advanced CORPORATE SOURCE:

Research Center, Kobe, 651-2492, Japan

SOURCE:

Biophysical Journal (2000), 78(6), 3048-3071

CODEN: BIOJAU; ISSN: 0006-3495 PUBLISHER: Biophysical Society

DOCUMENT TYPE: Journal LANGUAGE: English

AΒ Single-mol. and macroscopic reactions of fluorescent nucleotides with myosin have been compared. The single-mol. studies serve as paradigms for enzyme-catalyzed reactions and ligand-receptor interactions analyzed as individual stochastic processes. Fluorescent nucleotides, called Cy3-EDA-ATP and Cy5-EDA-ATP, were derived by coupling the dyes Cy3.29.0H and Cy5.29.0H with 2'(3')-O-[N-(2-aminoethyl) carbamoyl]ATP (EDA-ATP). The contraction of the coupling the dyes Cy3.29.0H and Cy5.29.0H with 2'(3')-O-[N-(2-aminoethyl) carbamoyl]ATP (EDA-ATP). ATP(ADP) analogs were sepd. into their resp. 2'- and 3'-O-isomers, the interconversion rate of which was 30[OH-] s-1 (0.016 h-1 at pH 7.1) at 22.degree.C. Macroscopic studies showed that 2'(3')-O-substituted nucleotides had properties similar to those of ATP and ADP in their interactions with myosin, actomyosin, and muscle fibers, although the ATP analogs did not relax muscle as well as ATP did. Significant differences in the fluorescence intensity of Cy3-nucleotide 2'- and 3'-O-isomers in free soln. and when they interacted with myosin were evident. Single-mol. studies using total internal reflection fluorescence microscopy showed that reciprocal mean lifetimes of the nucleotide analogs interacting with myosin filaments were one- to several-fold greater than predicted from macroscopic data. Kinetic and equil. data of nucleotide-(acto)myosin interactions derived from single-mol. microscopy now have a biochem. and physiol. framework. This is important for single-mol. mech. studies of motor proteins.

#### ΙT 288628-76-2P 288628-78-4P 288629-86-7P

RL: BPN (Biosynthetic preparation); BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (comparative single-mol. and ensemble myosin enzymol. using sulfoindocyanine ATP and ADP derivs.)

RN288628-76-2 HCAPLUS

CN Adenosine 5'-(trihydrogen diphosphate), 3'-[[2-[[6-[2-[3-[1-ethyl-1,3dihvdro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate] (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

\_so<sub>3</sub>-

RN 288628-78-4 HCAPLUS

CN Adenosine 5'-(trihydrogen diphosphate), 2'-[[2-[[6-[2-[3-[1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate] (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

## PAGE 1-A

PAGE 1-B

>so<sub>3</sub>-

HO3S

RN 288629-86-7 HCAPLUS

CN Adenosine 5'-(trihydrogen diphosphate), 2'(or 3')-[[2-[[6-[2-[5-[1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate] (9CI) (CA INDEX NAME)

CM 1

CRN 288629-84-5 CMF C36 H46 N4 O9 S2 CDES 8:EX

HO<sub>2</sub>C-NH-CH<sub>2</sub>-CH<sub>2</sub>-NH-C-(CH<sub>2</sub>)<sub>5</sub>

$$\begin{array}{c|c}
 & \text{Et} \\
 & \text{N}^{+} \\
 & \text{CH} \\
 & \text{Me} \\
 & \text{Me}$$

PAGE 1-B

∑so3H

CM 2

CRN 58-64-0

CMF C10 H15 N5 O10 P2

CDES 5:B-D-RIBO

Absolute stereochemistry.

## IT 192863-85-7P 288628-77-3P 288629-85-6P

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (comparative single-mol. and ensemble myosin enzymol. using sulfoindocyanine ATP and ADP derivs.)

RN 192863-85-7 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 3'-[[2-[[6-[2-[3-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

\_so3-

RN 288628-77-3 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 2'-[[2-[[6-[2-[3-[1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate] (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

## PAGE 1-A

PAGE 1-B

~so<sub>3</sub>-

PAGE 2-A

RN 288629-85-6 HCAPLUS CNethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-

pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-

oxohexyl]amino]ethyl]carbamate] (9CI) (CA INDEX NAME)

CM1

CRN 288629-84-5

CMF C36 H46 N4 O9 S2

CDES 8:EX

PAGE 1-B

─ SO3H

CM 2

CRN 56-65-5

CMF C10 H16 N5 O13 P3

CDES 5:B-D-RIBO

Absolute stereochemistry.

REFERENCE COUNT:

79 THERE ARE 79 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

### => d ibib abs hitstr 14

L14 ANSWER 14 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1999:511278 HCAPLUS

ACCESSION NUMBER: 1999:511278 HODOCUMENT NUMBER: 131:140472

TITLE: Dideoxy dye-labeled terminators for DNA sequencing

INVENTOR(S): Kumar, Shiv; Nampalli, Satyam; McArdle, Bernard F.;

Fuller, Carl W.

PATENT ASSIGNEE(S): Amersham Pharmacia Biotech, Inc., USA

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.				KIND		DATE			A	APPLICATION NO.				DATE			
	WO	WO 9940223 W: AU, CA,					19990812			W	WO 1999-US2104			4	19990202		<	
				BE,		CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,
		2319	777	02		-	1999			-			3197		1999			
	AU 9925717 EP 1060264			A1 A1		19990823 20001220					1999-25717 1999-90558			19990202 19990202				
			AT,		CH,	DE,	DK,	ES,	FR,						NL,			PT,
IE, FI JP 2002505853					Т2		20020226			J	P 20	00-5	3063	3	1999	0202	<	
PRIORITY APPLN. INFO.:										US 1	998-	1869	5	Α	1998	0204	<	
									1	WO 1	999-	US21	04	W	1999	0202	<	

OTHER SOURCE(S): MARPAT 131:140472

GI

AB A kit is provided for DNA sequencing comprising a first, second, third and fourth dye-labeled terminator mols., each of the dye terminator mols. comprising a dye mol., a linker of at least 10 atoms in length and a dideoxynucleoside mono- or triphosphate, and a thermostable DNA polymerase. The dye terminators provide uniform band intensities and the resoln. of dye-induced compression artifacts in DNA sequencing. The dideoxy dye-labeled terminators of the present invention are particularly well suited for use with DNA polymerases that are thermostable or which

Ι

contain an altered dNMP binding site. Their use do not require the use of nucleotide analogs such as dITP or .alpha.-thio nucleotides to eliminate dye-induced compression artifacts. There is a strong correlation between the length of the link between the dye mol. and the nucleotide and band uniformity, but little correlation between the type of dye (or other parameters) and band intensity. Dye terminators with linkers of 10 or more atoms up to 25 atoms when used in sequencing reactions produce bands in sequencing gels of significantly improved uniformity compared with dye terminators with linkers less than 10 atoms. In preferred embodiments, the dye terminators comprise structure I (B = 2',3'-dideoxy-7-deaza-ATP or -GTP or 2',3'-dideoxy-UTP or -CTP; L = linker attached to 7 position of purines or 5 position of pyrimidines; when B = deaza-ddATP or deaza-ddGTP, L = C.tplbond.CCH2NHCO(CH2)5; when B = ddUTP or ddCTP, L = C.tplbond.CCH2NHCO(CH2)5NHCO(CH2)5).

IT 235743-48-3P 235743-49-4P 235743-50-7P 235743-51-8P

RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses) (dideoxy dye-labeled terminators for DNA sequencing)

RN 235743-48-3 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[[6-[[3-[4-amino-1,2-dihydro-2-oxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-y1)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]-6-oxohexyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-B

RN 235743-49-4 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-1-[6-oxo-6-[[6-oxo-6-[[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-

tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]hexyl]amino]hexyl]-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

RN 235743-50-7 HCAPLUS

CN lH-Benz[e]indolium, 2-[5-[1,3-dihydro-1,1-dimethyl-3-[6-oxo-6-[[6-oxo-6-[[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]hexyl]amino]hexyl]-6,8-disulfo-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-B

RN 235743-51-8 HCAPLUS

1H-Benz[e]indolium, 2-[5-[3-[6-[[6-[[3-[4-amino-1,2-dihydro-2-oxo-1[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]-6oxohexyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-6,8-disulfo-2Hbenz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

5

REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

=> d ibib abs hitstr 15

L14 ANSWER 15 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1999:168972 HCAPLUS

DOCUMENT NUMBER: 131:2328

TITLE: Direct imaging of DNA in living cells reveals the

dynamics of chromosome formation

AUTHOR(S): Manders, Erik M. M.; Kimura, Hiroshi; Cook, Peter R.

CORPORATE SOURCE: Sir William Dunn School of Pathology, University of

Oxford, Oxford, OX1 3RE, UK

SOURCE: Journal of Cell Biology (1999), 144(5),

813-821

CODEN: JCLBA3; ISSN: 0021-9525 Rockefeller University Press

DOCUMENT TYPE: Journal LANGUAGE: English

Individual chromosomes are not directly visible within the interphase nuclei of most somatic cells; they can only be seen during mitosis. We have developed a method that allows DNA strands to be obsd. directly in living cells, and we use it to analyze how mitotic chromosomes form. A fluorescent analog (e.g., Cy5-dUTP) of the natural precursor, thymidine triphosphate, is introduced into cells, which are then grown on the heated stage of a confocal microscope. The analog is incorporated by the endogenous enzymes into DNA. As the mechanisms for recognizing and removing the unusual residues do not prevent subsequent progress around the cell cycle, the now fluorescent DNA strands can be followed as they assemble into chromosomes, and segregate to daughters and grand-daughters. Movies of such strands in living cells suggest that chromosome axes follow simple recognizable paths through their territories during G2 phase, and that late replicating regions maintain their relative positions as prophase chromosomes form. Quant. anal. confirms that individual regions move little during this stage of chromosome condensation. As a result, the gross structure of an interphase chromosome territory is directly related to that of the prophase chromosome.

IT 158613-49-1

PUBLISHER:

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses) (direct imaging of DNA in living cells reveals the dynamics of

(direct imaging of DNA in living cells reveals the dynamics of chromosome formation)

RN 158613-49-1 HCAPLUS

CN Uridine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[(1E)-3-[[6-[(2E)-2-[(2E,4E)-5-(3,3-dimethyl-5-sulfo-3H-indol-2-yl)-2,4-pentadienylidene]-2,3-dihydro-3,3-dimethyl-5-sulfo-1H-indol-1-yl]-1-oxohexyl]amino]-1-propenyl]-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

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PAGE 1-B

43

REFERENCE COUNT:

THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

### => d ibib abs hitstr 16

L14 ANSWER 16 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1998:597396 HCAPLUS

DOCUMENT NUMBER:

130:1959

TITLE:

New NIR dyes: synthesis, spectral properties and

applications in DNA analyses

AUTHOR(S):

Narayanan, Narasimhachari; Little, Garrick;

Raghavachari, Ramesh; Gibson, Jasmin; Lugade, Ananda;

Prescott, Chuck; Reiman, Kevin; Roemer, Steve; Steffens, Dave; Sutter, Scott; Draney, Daniel

CORPORATE SOURCE:

LI-COR, Inc., Biotech Division, Lincoln, NE, 68504,

USA

SOURCE:

NATO ASI Series, Series 3: High Technology (

1998), 52(Near-Infrared Dyes for High Technology Applications), 141-158 CODEN: NAHTF4; ISSN: 1383-7168

Kluwer Academic Publishers

PUBLISHER:

Journal English

DOCUMENT TYPE: LANGUAGE:

AB New pentamethine and heptamethine monofunctional asym. cyanine dyes have been synthesized. They are suitable for independently exciting at 680nm and 780nm laser diodes resp. The absorption and fluorescence characteristics such as molar absorptivity and quantum yield have been examd. in various solvents. A new spectrofluorometer, an instrument built in house is described. The dyes having a terminal hydroxyl group (1, 3, 4, 5 and 8) have been successfully attached to oligonucleotides on an automated DNA synthesizer through phosphoramidite chem. The dyes with carboxyl (2) and isothiocyanate functional groups (7) have been coupled directly to deoxyribonucleotides (dATP). The dye labeled primers and dye labeled dATPs provide excellent sensitivity and high throughput when used for sequencing and genotyping applications on LI-COR's 4200 automated DNA analyzer which independently detects at two wavelengths.

IT 215789-35-8P

RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses)

(synthesis, spectral properties of new NIR dyes and applications in DNA analyses)

RN 215789-35-8 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[6-[[9-[2-deoxy-5-O-[hydroxy[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-9H-purin-6-yl]amino]hexyl]amino]-6-oxohexyl]-1,3-dihydro-l,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfoobutyl)-, inner salt, tetralithium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

Me Me Me 
$$(CH_2)_{\overline{5}}$$
  $N_H$   $(CH_2)_{\overline{4}}$   $SO_3^-$ 

## PAGE 1-B

PAGE 2-A

PAGE 2-B

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REFERENCE COUNT:

30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

### => d ibib abs hitstr 17

L14 ANSWER 17 OF 26 HCAPLUS COPYRIGHT 2002 ACS 1998:514958 HCAPLUS ACCESSION NUMBER:

129:271927 DOCUMENT NUMBER:

ATPase kinetics of the Dictyostelium discoideum myosin TITLE:

II motor domain

Kuhlman, Philip A.; Bagshaw, Clive R. AUTHOR(S):

Department of Biochemistry, University of Leicester, CORPORATE SOURCE:

Leicester, LE1 7RH, UK
Journal of Muscle Research and Cell Motility ( SOURCE:

1998), 19(5), 491-504 CODEN: JMRMD3; ISSN: 0142-4319

PUBLISHER: Chapman & Hall

DOCUMENT TYPE: Journal LANGUAGE: English

AB Structural characterization of the mode of interaction of nucleotides bound to myosin has relied upon the crystal structure of the Dictyostelium discoideum myosin II motor domain. This fragment, denoted S1dC, lacks the regulatory domain and light chain subunits and may therefore fail to display the normal ATPase activity of the intact myosin mol. Here we show that the elementary steps of the Sldc ATPase pathway and the effects of actin are similar to those of the complete myosin head fragment. This indicates that truncation at residue E759, with the removal of the light chain binding sites, is not crucial to catalytic activity. In particular, SldC does not show the anomalous tight binding of ADP displayed by the slightly shorter M754 construct reported elsewhere. We also show that the fluorescent analog Cy3-EDA-ATP is a good substrate for SldC and demonstrate the use of fluorescence correlation spectroscopy to det. the affinity of Cy3-EDA-ADP using microgram quantities of proteins.

213904-25-7 213904-27-9 TΨ

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(ATPase kinetics of the Dictyostelium discoideum myosin II motor domain)

RN 213904-25-7 HCAPLUS

Adenosine 5'-(tetrahydrogen triphosphate), 2'-[[2-[[6-[2-[3-[1-[6-[(2,5-CNdioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2Hindol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

PAGE 2-A

RN 213904-27-9 HCAPLUS

CN Adenosine 5'-(tetrahydrogen triphosphate), 3'-[[2-[[6-[2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-

indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

PAGE 1-A

PAGE 1-B

~so3-

## MAUPIN 09/829,467

### => d ibib abs hitstr 18

L14 ANSWER 18 OF 26 HCAPLUS COPYRIGHT 2002 ACS

1998:312612 HCAPLUS ACCESSION NUMBER:

129:78684 DOCUMENT NUMBER:

Time-resolved identification of individual TITLE:

mononucleotide molecules in aqueous solution with

pulsed semiconductor lasers

Sauer, Markus; Arden-Jacob, Jutta; Drexhage, Karl H.; Gobel, Florian; Lieberwirth, Ulrike; Muhlegger, Klaus; AUTHOR(S):

Muller, Ralph; Wolfrum, Jurgen; Zander, Christoph

CORPORATE SOURCE: Physikalisch-Chemisches Institut, Universitat

Heidelberg, Heidelberg, 69120, Germany Bioimaging (1998), 6(1), 14-24

SOURCE:

CODEN: BOIMEL; ISSN: 0966-9051 Institute of Physics Publishing

PUBLISHER: DOCUMENT TYPE: Journal LANGUAGE: English

We applied a short-pulse diode laser emitting at 640 nm with a repetition rate of 56 MHz in combination with a confocal microscope to study bursts of fluorescence photons from individual differently labeled mononucleotide mols. in water. Two newly synthesized dyes, an oxazine dye (MR121) and a rhodamine dye (JA53), and two com. available dyes, a carbocyanine dye (Cy5) and a bora-diaza-indacene dye (Bodipy630/650), were used as fluorescent labels. The time-resolved fluorescence signals of individual mononucleotide mols. in water were analyzed and identified by a max. likelihood estimator (MLE). Taking only those single mol. transits which contain more than 30 collected photoelectrons, the two labeled mononucleotide mols., Cy5-dCTP and Bodipy-dUTP, can be identified by time-resolved fluorescence spectroscopy with a probability of correct classification of greater than 99%. Our results show that at least three differently labeled mononucleotide mols. can be identified in a common aq. soln. We obtain an overall classification probability of 90% for the time-resolved identification of Cy5-dCTP, MR121-dUTP and Bodipy-dUTP mols. via their characteristic fluorescence lifetimes of 1.05.+-.0.33 ns (Cy5-dCTP), 2.07.+-.0.59 ns (MR121-dUTP) and 3.88.+-.1.71 ns (Bodipy-dUTP).

#### 206271-55-8 TΤ

RL: ANT (Analyte); ANST (Analytical study) (time-resolved identification of individual mononucleotide mols. in aq. soln. with pulsed semiconductor lasers)

RN 206271-55-8 HCAPLUS

Cytidine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[2-[(1E,3E,5E)-CN 5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1propynyl]-, inner salt, ion(5-) (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry as shown.

-03S

# PAGE 1-B

### => d ibib abs hitstr 19

L14 ANSWER 19 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:220741 HCAPLUS

DOCUMENT NUMBER: 128:303476

TITLE: Single-molecule counting and identification in a

microcapillary

AUTHOR(S): Zander, C.; Drexhage, K. H.; Han, K.-T.; Wolfrum, J.;

Sauer, M.

CORPORATE SOURCE: Im Neuenheimer Feld 253, Physikalisch-Chemisches

Institut, Universitat Heidelberg, Heidelberg, D-69120,

Germany

SOURCE: Chemical Physics Letters (1998), 286(5,6),

457-465

CODEN: CHPLBC; ISSN: 0009-2614

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Using a confocal microscope the authors studied photon bursts from individual mols. (dye-labeled mononucleotides) flowing in a cone-shaped microcapillary with an inner diam. of 0.5 .mu.m at the small end of the cone. The flow of the conjugates was established by electrokinetic forces. Excitation of the fluorophore was provided by a pulsed diode laser (.lambda. = 640 nm, av. power 800 .mu.W, repetition rate 56 MHz). The characteristic diffusion and flow time through the laser focus and burst size statistics were detd. in the microcapillary as well as in an open vol. Applying time-correlated single-photon counting, two different conjugate species (Cy5-dCTP, JA53-dUTP) can be distinguished due to their characteristic fluorescence decay time with a probability of correct classification of 80%.

IT 206271-55-8, Cy 5dCTP tetraanion

RL: ANT (Analyte); ANST (Analytical study)

(Cy 5dCTP; single-mol. counting and identification in microcapillary)

RN 206271-55-8 HCAPLUS

CN Cytidine 5'-(tetrahydrogen triphosphate), 2'-deoxy-5-[3-[[6-[2-[(1E,3E,5E)-5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]-1-propynyl]-, inner salt, ion(5-) (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

-03S

PAGE 1-B

### => d ibib abs hitstr 20

L14 ANSWER 20 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:71282 HCAPLUS

128:138318

DOCUMENT NUMBER: TITLE:

Sensors

INVENTOR(S):

Issachar, David

PATENT ASSIGNEE(S):

Sensors Technology Company B.V., Neth.; Issachar,

David

SOURCE:

PCT Int. Appl., 31 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                    KIND DATE
                                                   APPLICATION NO. DATE
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                                                   ______
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                                             WO 1997-IL221 19970702 <--
                                 19980122
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               DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,
          PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
               GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
               GN, ML, MR, NE, SN, TD, TG
                         A1 19980209
                                                   AU 1997-32714
                                                                        19970702 <--
     AU 9732714
PRIORITY APPLN. INFO.:
                                                                        19960715 <--
                                                IL 1996-118859
                                                WO 1997-IL221
                                                                        19970702 <--
     The present invention concerns a sensor for detection of a biol. analyte
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- AB The present invention concerns a sensor for detection of a biol. analyte in a test sample. The biol. analyte is a member of a pair forming group such as antigen/antibody, ligand/receptor, etc. The sensor comprises a porous matrix where in each cavity are entrapped a mol. capable of specifically binding to the analyte as well as an analyte-analog. Competitive displacement of the analyte-analog by the assayed analyte bridge at least one detectable property of the sensor.
- IT 202413-97-6P
  - RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses) (sensor for detection of a biol. analyte in a test sample)
- RN 202413-97-6 HCAPLUS
- CN Poly(oxy-1,2-ethanediyl), .alpha.-(2-aminoethyl)-.omega.-[2-[[(2S)-6-[[[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]carbonyl]amino]-1-oxo-2-[[(1,2,3,6-tetrahydro-1,3-dimethyl-2,6-dioxo-7H-purin-7-yl)acetyl]amino]hexyl]amino]ethoxy]-, inner salt (9CI) (CA INDEX NAME)

PAGE 2-A

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IT 202414-00-4D, antibody conjugate

RL: RCT (Reactant); RACT (Reactant or reagent) (sensor for detection of a biol. analyte in a test sample)

RN 202414-00-4 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-[2-[[4-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)benzoyl]amino]ethyl]-.omega.-[2-[[(2S)-6-[[[6-[2-[5-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]carbonyl]amino]-1-oxo-2-[[(1,2,3,6-tetrahydro-1,3-dimethyl-2,6-dioxo-7H-purin-7-yl)acetyl]amino]hexyl]amino]ethoxy]-, inner salt (9CI) (CA INDEX NAME)

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 & O \\
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 & O \\
 & C \\
 & O \\$$

HO3S

PAGE 1-B

PAGE 2-B

| Me

## MAUPIN 09/829,467

## => d ibib abs hitstr 21

L14 ANSWER 21 OF 26 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1997:472614 HCAPLUS

DOCUMENT NUMBER: 127:145698

DOCUMENT NUMBER: 127:145696

TITLE: Molecular mechanism controlling the incorporation of

fluorescent nucleotides into DNA by PCR

AUTHOR(S): Zhu, Zhengrong; Waggoner, Alan S.

CORPORATE SOURCE: Center for Light Microscope Imaging and Biotechnology,

Carnegie Mellon University, Pittsburgh, PA, USA

SOURCE: Cytometry (1997), 28(3), 206-211

CODEN: CYTODQ; ISSN: 0196-4763

PUBLISHER: Wiley-Liss
DOCUMENT TYPE: Journal
LANGUAGE: English

AB The efficiency and yield of incorporation of fluorescent nucleotides into DNA by polymerase chain reaction (PCR) have been investigated with linear amplification (PCR with single-stranded template and single primer). In the present study, we prepd. single-stranded templates with defined sequences and used dUTP attached to the fluorescent label with linkers of different lengths. Incorporation and yield of the modified dUTP were reduced when the sequence demanded that multiple dyes be inserted at adjacent sites. The interactions between the polymerase and cyanine-labeled sites on the extending strand probably terminated the chain extension. Thus, because labeling d. was increased, the yield of PCR was reduced. We also found that the interactions between the primer and dye-labeled sites on template disturb primer annealing and lead to a decrease in PCR yield.

IT 159018-62-9 159018-64-1

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(mol. mechanism controlling the incorporation of fluorescent nucleotides into DNA by PCR)

RN 159018-62-9 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[6-[[3-[1-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-2-propenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

RN 159018-64-1 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[6-[[6-[[6-[[3-[1-[2-deoxy-5-0-[hydroxy[hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-2-propenyl]amino]-6-oxohexyl]amino]-6-oxohexyl]amino]-6-oxohexyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

# PAGE 1-B

#### MAUPIN 09/829,467

#### => d ibib abs hitstr 22

L14 ANSWER 22 OF 26 HCAPLUS COPYRIGHT 2002 ACS 1997:390729 HCAPLUS

ACCESSION NUMBER:

127:119183 DOCUMENT NUMBER:

Single molecule imaging of fluorophores and enzymic TITLE:

reactions achieved by objective-type total internal

reflection fluorescence microscopy

Tokunaga, Makio; Kitamura, Kazuo; Saito, Kiwamu; AUTHOR(S):

Iwane, Atsuko Hikikoshi; Yanagida, Toshio

Yanagida BioMotron Project, ERATO, JST, Osaka, 562, CORPORATE SOURCE:

Japan

Biochemical and Biophysical Research Communications ( SOURCE:

**1997**), 235(1), 47-53 CODEN: BBRCA9; ISSN: 0006-291X

PUBLISHER: Academic DOCUMENT TYPE: Journal LANGUAGE: English

Imaging of single fluorescence mols. has been achieved in a relatively simple manner using objective-type total internal reflection fluorescence microscopy (TIRFM). Switching from epi-fluorescence microscopy to objective-type TIRFM was achieved by translation of a single mirror in the system. Clear images of single mols. of an orange fluorescent dye, Cy3, were obtained with a fluorescence-to-background ratio of 12, using a conventional high aperture objective (PlanApo, 100 .times., Na 1.4) with ordinary coverslips and immersion oil. This method allowed visualization of single mols. under scanning probe microscopes. Taking advantage of the technique of single mol. imaging, individual ATP turnovers have been visualized with a fluorescent ATP analog, Cy3-ATP, using a simple exptl. strategy. Clear on/off signals were obtained that correspond to the assocn. and dissocn. of single Cy3-ATP/ADP mols. with a single myosin head This method will allow a variety of single-mol. assays of biomol. functions to be performed using fluorescently labeled substrates, ligands, messengers, and biol. active mols. Thus, the present technique provides a simple yet powerful and universal tool for researchers to probe the events of single mols.

#### IT 192863-85-7

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (single mol. imaging of fluorophores and enzymic reactions by total internal reflection fluorescence microscopy)

192863-85-7 HCAPLUS RN

CN dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1-propenyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]ethyl]carbamate], inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

~so3-

PAGE 2-A

#### => d ibib abs hitstr 23

L14 ANSWER 23 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1996:572057 HCAPLUS

DOCUMENT NUMBER:

125:214232

TITLE:

Stabilization of labeled nucleoside triphosphates with

magnesium-binding compounds

INVENTOR(S):

Duthie, R. Scott; Brush, Charles K.; Stirchak, Eugene

P.; Freeman, Mark E.; Burazin, Lawrence J.

PATENT ASSIGNEE(S):

Pharmacia Biotech Inc., USA

SOURCE:

PCT Int. Appl., 23 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent English

LANGUAGE:

m. 1

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PA'	rent	NO.		KI	ND	DATE			A	PP:	LICA	)ITA	N NC	Ο.	DATE			
WO	9622	<del></del> -		 A	 1	1996	0725		– W	0	1996	5-US	- <b></b> 5274		1996	0105	<	
	W:	ΑU,	CA,	JΡ														
	RW:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	G.	R, ]	[Ε,	ΙT,	LU,	MC,	NL,	PT,	SE
US	5808	043		Α		1998	0915		U	S	1995	5-37	7445	6	1995	0118		
CA	2210	900		A	Ą	1996	0725		С	Ά	1996	5-22	2109	00	1996	0105	<	
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JP	1050	4974		T	2	1998	0519		J	Ρ	1996	6-52	2230	3	1996	0105	<	
JP	3093	275		B:	2	2000	1003											
PRIORIT	Y APP	LN.	INFO	. :				1	US 1	99	5-37	7445	56	A	1995	0118	<	
								1	WO 1	99	6-US	3274	4	W	1996	0105	<	

AB A prepn. of a labeled nucleotide comprising at least one compd. having a Mg2+ assocn. const. between 1 .times. 10-11 to 1 .times. 10-2, inclusive, is claimed. The compd. is preferably selected from the group consisting of citrate, isocitrate, phosphate, EGTA, EDTA, and EDTA. The concn. of the compd. is preferably at least 5 mM.

IT 174817-56-2

RL: MSC (Miscellaneous)

(stabilization of labeled nucleoside triphosphates with magnesium-binding compds.)

RN 174817-56-2 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[[9-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonoox y)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-9H-purin-6-yl]amino]hexyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

# PAGE 1-B

#### MAUPIN 09/829,467

#### => d ibib abs hitstr 24

L14 ANSWER 24 OF 26 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:126958 HCAPLUS

DOCUMENT NUMBER: 124:224660

TITLE: Characterization of fluorescent nucleoside

triphosphates by capillary electrophoresis with laser-induced fluorescence detection: action of

alkaline phosphatase and DNA polymerase

AUTHOR(S): Evangelista, Ramon A.; Liu, Ming-Sun; Rampal, Sushma;

Chen, Fu-Tai A.

CORPORATE SOURCE: Advanced Technology Center, Beckman Instruments Inc.,

Fullerton, CA, 92634, USA

SOURCE: Analytical Biochemistry (1996), 235(1),

89-97

CODEN: ANBCA2; ISSN: 0003-2697

PUBLISHER: Academic
DOCUMENT TYPE: Journal
LANGUAGE: English

AB A method of anal. of fluor-labeled nucleoside triphosphates based on alk. phosphatase-catalyzed sequential cleavage of phosphate groups with monitoring of all fluorescent species by capillary electrophoresis with laser-induced fluorescence detection is presented. The method allows detn. of the purity of the triphosphate samples as well as the relative amts. of the lower phosphate contaminants. The ability of one of the fluor-labeled nucleoside triphosphates to serve as polymerase substrate was verified by labeling DNA restriction fragments by the method of filling recessed 3'-ends using DNA polymerase Klenow fragment.

IT 174817-57-3

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(fluorescent nucleoside triphosphates anal. by capillary electrophoresis with laser-induced fluorescence and action of enzymes)

RN 174817-57-3 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[[6-[[1-[2-deoxy-5-O-[hydroxy[[hydroxy(phosphonoox y)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2-dihydro-2-oxo-4-pyrimidinyl]amino]hexyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

generic search

#### MAUPIN 09/829,467

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L9
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                85-32-5/BI OR 86-01-1/BI OR 95-50-1/BI)
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                                                L10 AND L11
L15
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                                                 ?STYRYL?
L16
         863910 SEA FILE=HCAPLUS ABB=ON
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                NUCLEIC)
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L19
           6154 SEA FILE=HCAPLUS ABB=ON PLU=ON "PHOSPHATES, BIOLOGICAL
                STUDIES"/CT
L34
                STR
                             any cydo RING
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                            10 @11 12
                        C-chain of De 2-9 carbons
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REP G2=(2-9) C
NODE ATTRIBUTES:
CONNECT IS E3 RC AT
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DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 14

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STEREO ATTRIBUTES: NONE
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L40	639	SEA	FILE=REGISTRY	ABB=ON	PLU=ON	L36 AND ?PHOSPH?/CNS
1.41	39918	SEA	FILE=REGISTRY	ABB=ON	PIJU=ON	L36 NOT (L39 OR L40)

# MAUPIN 09/829,467

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L44	261	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L40 DNA ST
L45	219	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L40 DNA, ETC
L46	8330	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L16 (3A) FLUORES?
L47	33	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L46 AND (L43 OR L44)
L48	9	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L47 AND ?CONJUGAT?
L49	24	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L47 NOT L48
L50	68099	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L15 OR ?CYANINE?
L51	16	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L49 AND L50
L53	7	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	(?SULFON? OR ?PHOSPH?) AND
		L51			
L55	52	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L45 AND ((L18 OR L19) OR
		?SU	LFON? OR ?PHOSPH?)		
L56	1	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L55 AND L16
L58	2	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L16 AND L45
L59	2	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L58 OR L56
L60	17	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	(L48 OR L53 OR L59) NOT L12

17 cites

#### MAUPIN 09/829,467

#### => d ibib abs hitstr 160 1

L60 ANSWER 1 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:787923 HCAPLUS

136:398000

TITLE:

Single- and dual- near-infrared fluorescent

labeled nucleic acid conjugate for

nucleic acid detection

AUTHOR(S):

Lin, Zhihong; Wu, Meng; Ren, Shu; Arbter, Michaela; Boehmer, Martin; Mirsky, Vladimir; Wolfbeis, Otto S. Department of Chemistry, Tongji Medical College,

CORPORATE SOURCE:

Huazhong University of Science and Technology, Wuhan,

Hubei, 430030, Peop. Rep. China

SOURCE:

Proceedings of SPIE-The International Society for Optical Engineering (2001), 4414(International Conference on Sensor Technology (ISTC 2001), 2001),

111-114

CODEN: PSISDG; ISSN: 0277-786X

PUBLISHER:

SPIE-The International Society for Optical Engineering

DOCUMENT TYPE: Journal LANGUAGE: English

In this research, a near-IR fluorescent labeled nucleic acid conjugate for the nucleic acid detection was synthesized, and characterized preliminarily for the detection of the nucleic acid. The **conjugate** combines the mol. recognition properties of the oligonucleotides with the near-IR fluorescence label PR 646. Both singleand dual- labeled conjugates were studied for their hybridization with the complementary nucleic acid. The dual labeled conjugate has indicated that the self-quenching effect exists in

ssDNA form while the fluorescence increases greatly after hybridization with the complementary nucleic acids. The time-resolved fluorescence was also studied.

IT 429681-54-9, PR 646

> RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (single- and dual- near-IR fluorescent labeled nucleic acid conjugate for nucleic acid detection)

RN 429681-54-9 HCAPLUS

CN (ethoxyhydroxyphosphinyl)ethyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

#### => d ibib abs hitstr 160 2

L60 ANSWER 2 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:507799 HCAPLUS

DOCUMENT NUMBER:

135:93921

TITLE:

Mobility-modifying cyanine dyes

INVENTOR(S):

Menchen, Steven M.; Benson, Scott C.; Rosenblum,

Barnett B.; Khan, Shaheer H.

PATENT ASSIGNEE(S):

PE Corporation, USA

SOURCE:

PCT Int. Appl., 133 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

Engli

PATENT INFORMATION:

PATENT NO. KIND			DATE			APPLICATION NO.					DATE						
							_										
WO 2001049790		A.	2	20010712			WO 2001-US152 20010103										
WO :	2001	0497	90	A.	3	2001	1206										
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,
		HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	ΝZ,	PL,	PT,	RO,	RU,
		SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	UZ,	VN,	YU,
		ZA,	ZW,	AM,	ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM					
	RW:	GH,	GM,	KE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,
		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG		
RITY	APP	LN.	INFO	. :				i	US 2	000-	4772	70	Α	2000	0104		

OTHER SOURCE(S): MARPAT 135:93921

The present invention provides a novel class of fluorescent cyanine dye compds. that are modified at one of the heterocyclic ring nitrogen atoms with a mobility-modifying moiety that permits the electrophoretic mobilities of polynucleotides labeled with the mobility-modifying cyanine dyes to be adjusted or tuned in a predictable fashion while retaining enzymic activity. The ability to predictably tune the relative electrophoretic mobilities of the dyes permits the creation of sets of mobility-matched fluorescent dyes of a variety of structures for a variety of applications, including fluorescence-based 4-color nucleic acid sequencing reactions.

IT 349491-68-5P 349491-69-6P 349491-73-2P 349491-74-3P 349491-76-5P 349491-78-7P

RL: ARG (Analytical reagent use); IMF (Industrial manufacture); TEM (Technical or engineered material use); ANST (Analytical study); PREP (Preparation); USES (Uses)

(mobility-modifying fluorescent cyanine dyes for nucleic acid sequencing reactions)

RN 349491-68-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[3-[4-amino-7-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-7H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propynyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-[(2,4-disulfophenyl)methyl]-1,1-dimethyl-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

## PAGE 1-B

PAGE 2-A

RN 349491-69-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[2-[[3-[4-amino-7-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-7H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propynyl]oxy]ethyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

PAGE 1-A

Me Me Me 
$$(CH_2)_{\overline{5}}$$
  $N_H^+$   $(CH_2)_{\overline{3}}$   $SO_3^-$ 

PAGE 1-B

// \

PAGE 2-B

RN 349491-73-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[3-[2-amino-4,7-dihydro-4-oxo-7-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-1H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propynyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-[(2,4-disulfophenyl)methyl]-1,1-dimethyl-,inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

PAGE 1-B

PAGE 2-A

RN 349491-74-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[2-[[3-[2-amino-4,7-dihydro-4-oxo-7-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-1H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propynyl]oxy]ethyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

PAGE 1-A

Me Me Me Me 
$$(CH_2)_{\overline{5}}$$
  $(CH_2)_{\overline{5}}$   $(CH_2)_{\overline{5}}$ 

PAGE 1-B

RN 349491-76-5 HCAPLUS
CN 1H-Benz[e]indolium, 2-[5-[3-[6-[[3-[4-amino-1,2-dihydro-2-oxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]-6-oxohexyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-[(2,4-disulfophenyl)methyl]-1,1-dimethyl-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

PAGE 1-B

PAGE 2-A

SO3H

RN 349491-78-7 HCAPLUS CN 1H-Benz[e]indolium, 2

## (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

## PAGE 1-B

#### => d ibib abs hitstr 160 3

L60 ANSWER 3 OF 17 HCAPLUS COPYRIGHT 2002 ACS 2001:440613 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

136:145683

TITLE:

Novel fluorescent nonnucleoside triphosphates as terminators of enzyme-directed DNA synthesis

AUTHOR(S):

Roemer, Stephen C.; Johnson, Craig M.; Boveia, Vince

R.; Buzby, Philip R.; DiMeo, James J.; Draney, Dan; Narayanan, Narasimhachari; Olive, D. Michael

CORPORATE SOURCE:

Orchid BioSciences, Inc., Princeton, NJ, USA

SOURCE:

Proceedings of SPIE-The International Society for Optical Engineering (2001), 4264 (Genomics and

Proteomics Technologies), 1-8 CODEN: PSISDG; ISSN: 0277-786X

PUBLISHER:

SPIE-The International Society for Optical Engineering

DOCUMENT TYPE:

Journal

LANGUAGE: English

Near IR (NIR) fluorescent acycloterminators were tested as substrates in the Sanger enzymic method of DNA sequencing. The acyclic triphosphates of adenosine, uridine, guanosine, and cytidine (AcyNTP) were labeled with a heptamethine carbocyanine dye via a propargylamino linker to the purine or pyrimidine base. Dye-labeled AcyNTPs which are lacking in the sugar moiety positions equiv. to the C-2 and C-3 of the ribose functioned similarly to chain-terminating dideoxynucleotides (ddNTPs). These fluorescent nonnucleotide analogs were incorporated by a mutant, thermostable Taq DNA polymerase with the same efficacy and fidelity as traditional ddNTPs. Sequence read length and base-calling accuracy were comparable for both dye- acycloterminator and dye-primer sequencing methods. In two primer walking projects, cycle sequencing with fluorescent AcyNTPs achieved a mean sequence read length of 1,090 bases with 99.1% accuracy at one kilobase read length. The cyanine dye-labeled acycloterminators produced electropherograms in which weak T peaks follow G peaks. In cases of polymorphism, such peak height variability may make it difficult to distinguish the presence or absence of a heterozygote at a specific site.

395645-36-0 395645-37-1 395645-38-2

395645-39-3

RL: ARG (Analytical reagent use); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses) (novel fluorescent nonnucleoside triphosphates as terminators of enzyme-directed DNA synthesis)

395645-36-0 HCAPLUS RN

CN 3H-Indolium, 2-[2-[2-[4-[3-[3-[4-amino-7-(6,8,10,10-tetrahydroxy-6,8,10trioxido-2,5,7,9-tetraoxa-6,8,10-triphosphadec-1-yl)-7H-pyrrolo[2,3d]pyrimidin-5-yl]-2-propynyl]amino]-3-oxopropyl]phenoxy]-3-[2-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene}ethyl]-1-cyclohexen-1yl]ethenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

#### PAGE 1-B

## Na

RN 395645-37-1 HCAPLUS

CN 3H-Indolium, 2-[2-[2-[4-[3-[[3-[2-amino-4,7-dihydro-4-oxo-7-(6,8,10,10-tetrahydroxy-6,8,10-trioxido-2,5,7,9-tetraoxa-6,8,10-triphosphadec-1-yl)-1H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propynyl]amino]-3-oxopropyl]phenoxy]-3-[2-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethyl]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

## PAGE 1-B

#### Na

RN 395645-38-2 HCAPLUS
CN 3H-Indolium, 2-[2-[3-[2-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethyl]-2-[4-[3-oxo-3-[[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-(6,8,10,10-tetrahydroxy-6,8,10-trioxido-2,5,7,9-tetraoxa-6,8,10-triphosphadec-1-yl)-5-pyrimidinyl]-2-propynyl]amino]propyl]phenoxy]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

# PAGE 2-A

## Na

## RN 395645-39-3 HCAPLUS

CN 3H-Indolium, 2-[2-[4-[3-[[3-[4-amino-1,2-dihydro-2-oxo-1-(6,8,10,10-tetrahydroxy-6,8,10-trioxido-2,5,7,9-tetraoxa-6,8,10-triphosphadec-1-yl)-5-pyrimidinyl]-2-propynyl]amino]-3-oxopropyl]phenoxy]-3-[2-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethyl]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 2-A

Na

REFERENCE COUNT:

25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

#### => d ibib abs hitstr 160 4

L60 ANSWER 4 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:310508 HCAPLUS

DOCUMENT NUMBER: 134:323136

TITLE: Cyanine dyes as labeling reagents for detection of

biological and other materials by luminescence methods

INVENTOR(S): Waggoner, Alan S.

PATENT ASSIGNEE(S): Carnegie Mellon University, USA

SOURCE: U.S., 20 pp., Cont.-in-part of U.S. 5,627,027.

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO. DATE
US 6225050	B1	20010501	US 1996-745712 19961112
US 5268486	Α	19931207	US 1992-884636 19920515
US 5627027	Α	19970506	US 1992-831759 19920922
US 5486616	A	19960123	US 1993-158952 19931129
US 5569766	Α	19961029	US 1993-158953 19931129
US 5569587	Α	19961029	US 1995-424219 19950419
US 6048982	Α	20000411	US 1997-873470 19970612
PRIORITY APPLN. INFO.	:		US 1986-854347 B1 19860418
			US 1992-831759 A2 19920922
			US 1988-240756 B1 19880902
•			US 1992-882802 B1 19920514
			US 1992-884636 A3 19920515
			US 1996-745712 A3 19961112

OTHER SOURCE(S): MARPAT 134:323136

Cyanine and related dyes, such as merocyanine, styryl and oxonol dyes, are strongly light-absorbing and highly luminescent. Cyanine and related dyes having functional groups to make them reactive with amine, hydroxy and sulfhydryl groups are covalently attached to proteins, nucleic acids, carbohydrates, sugars, cells and combinations thereof, and other biol. and nonbiol. materials, to make these materials fluorescent so that they can be detected. The labeled materials can then be used in assays employing excitation light sources and luminescence detectors. For example, fluorescent cyanine and related dyes can be attached to amine, hydroxy or sulfhydryl groups of avidin and to antibodies and to lectins. Thereupon, avidin labeled with cyanine type dyes can be used to quantify biotinylated materials and antibodies conjugated with cyanine-type dyes can be used to detect and measure antigens and haptens. In addn., cyanineconjugated lectins can be used to detect specific carbohydrate groups. Also, cyanine-conjugated fragments of DNA or RNA can be used to identify the presence of complementary nucleotide sequences in DNA or RNA.

# IT 336850-25-0DP, nucleotide conjugate 336850-27-2DP, nucleotide conjugate

RL: NUU (Other use, unclassified); PRP (Properties); SPN (Synthetic preparation); PREP (Preparation); USES (Uses)

(as **fluorescent** primer; cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN 336850-25-0 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[5-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]pentyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI)

(CA INDEX NAME)

PAGE 1-A

N(Pr-i)2

(CH2)5-0-P-0-CH2

N+

CH=CH-CH=CH-CH

Me

Me

Me

PAGE 1-B

— cн<sub>2</sub>- сn

RN 336850-27-2 HCAPLUS
CN 1H-Benz[e]indolium, 2-[5-[3-[5-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]pentyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo-, inner salt, potassium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Et

O3S

CH—CH—CH—CH—CH—(CH<sub>2</sub>)5-O-P-O-CH<sub>2</sub>

Me

N(Pr-i)2

Me

N

Me

N

Me

N

Me

K

PAGE 1-B

— сн<sub>2</sub>— си

#### 336850-25-0P IT

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(as reactive dye; cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN

336850-25-0 HCAPLUS 3H-Indolium, 2-[5-[1-[5-[[[bis(1-methylethyl)amino](2-CN cyanoethoxy)phosphino]oxy]pentyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A N(Pr-i)2 (CH2) 5-O-P-O-CH2 CH CH CH CH CH Me Me -03S Мe Me

PAGE 1-B

— сн<sub>2</sub>- си

#### IT 336850-30-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN 336850-30-7 HCAPLUS

CN 1H-Benz[e] indolium, 2-[5-[3-[5-[[[bis(1-methylethyl)amino](2-methylethyl)amino])]cyanoethoxy)phosphino]oxy]pentyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- CH2-CN

IT 336850-27-2P

RL: SPN (Synthetic preparation); PREP (Preparation) (cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN 336850-27-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[5-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]pentyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo-, inner salt, potassium salt (9CI) (CA INDEX NAME)

PAGE 1-A

K

PAGE 1-B

— сн<sub>2</sub>— си

REFERENCE COUNT:

19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

#### MAUPIN 09/829,467

=> d ibib abs hitstr 160 5

L60 ANSWER 5 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:178461 HCAPLUS

DOCUMENT NUMBER: 134:217985

TITLE: Method for 3' end-labeling ribonucleic acids

INVENTOR(S): Ach, Robert A.

PATENT ASSIGNEE(S): Agilent Technologies Inc., USA

SOURCE: U.S., 8 pp.
CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO. DATE	
US 6201112	B1	20010313	US 1999-359564 19990722	
US 2001009762	A1	20010726	US 2001-802358 20010309	
PRIORITY APPLN. INFO.:			US 1999-359564 A3 19990722	

AB Methods of end-labeling ribonucleic acids with non-radioactively labeled ribonucleotides, and particularly fluorescently labeled ribonucleotides, are provided. In the subject methods, a RNA is contacted with a non-radioactively labeled ribonucleotide in the presence of a prokaryotic, usually bacterial, poly(A) polymerase under conditions sufficient for covalent bonding of the labeled ribonucleotide to the 3' end of the RNA to occur. Also provided are kits for practicing the subject method. The subject methods and kits find use in a variety of applications where labeling of the 3' end of a RNA with a non-radioactive label, particularly a fluorescent label, is desired.

IT 329320-43-6

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (method for 3' end-labeling ribonucleic acids)

RN 329320-43-6 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[6-[[3-[4-amino-7-[5-O-[hydroxy[[hydroxy(phosphonooxy )phosphinyl]oxy]phosphinyl]-.beta.-D-ribofuranosyl]-7H-pyrrolo[2,3-d]pyrimidin-5-yl]-2-propenyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-B

REFERENCE COUNT:

13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d ibib abs hitstr 160 6

L60 ANSWER 6 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:93900 HCAPLUS

DOCUMENT NUMBER: 134:164473

TITLE: Acylsulfonamido-substituted polymethine

fluorescent dyes and their use as fluorescent coloring

materials and/or markers for biomolecules

Deroover, Geert; Missfeldt, Michael; Simon, Lydia INVENTOR(S):

PATENT ASSIGNEE(S): Bayer A.-G., Germany SOURCE: Ger. Offen., 68 pp. CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
KIND DATE APPLICATION NO. DATE
          PATENT NO.
                                    KIND DATE
         DE 19937024 A1 20010208 DE 1999-19937024 19990805
WO 2001011370 A1 20010215 WO 2000-EP7070 20000724
                         AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
          EP 1206703
                                            A1 20020522 EP 2000-958289 20000724
                  R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL
PRIORITY APPLN. INFO.:
                                                                                   DE 1999-19937024 A 19990805
                                                                                   WO 2000-EP7070 W 20000724
```

MARPAT 134:164473 OTHER SOURCE(S):

Polymethine dyes contg. (1) at least one acylsulfonamido group of the formula (CH2)nYNHAR, where A and Y are electron-donating groups such as CO or SO2, R = optionally substituted alkyl or aryl, and n = 1-9and (2) and at least one other functional group are effective as fluorescent coloring materials or markers for biomols. The polymethine dyes have improved light stability compared to prior-art indole or squaric acid-based materials when used with RNA, DNA, or proteins. Examples of prepn. of 2 dyes were given.

324745-35-9 324745-37-1 ΙT

> RL: BUU (Biological use, unclassified); TEM (Technical or engineered material use); BIOL (Biological study); USES (Uses)

(acylsulfonamido-substituted polymethine fluorescent dye markers for biomols.)

RN 324745-35-9 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-1-[2-[(methylsulfonyl)amino]-2oxoethyl]-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-[6-[(2,5-dioxo-1pyrrolidinyl)oxy]-6-oxohexyl]-3,3-dimethyl-5-sulfo-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

## PAGE 2-A

## K

RN 324745-37-1 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[1,3-dihydro-1,1-dimethyl-3-[2-[(methylsulfonyl)amino]-2-oxoethyl]-7-sulfo-2H-benz[e]indol-2-ylidene]-1,3pentadienyl]-3-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,1-dimethyl-7-sulfo-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

K

#### => d ibib abs hitstr 160 7

L60 ANSWER 7 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:10685 HCAPLUS

DOCUMENT NUMBER: 134:102214

TITLE: New fluorescent cyanine labels containing a

sulfonamido linker arm

INVENTOR(S): Caputo, Giuseppe; Della, Ciana Leopoldo

PATENT ASSIGNEE(S): Innosense S.r.L., Italy SOURCE: Eur. Pat. Appl., 94 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

LANGUAGE: Eng FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

EP 1065250 Al 20010103 EP 1999-112696 19990702

R: AT BE CH DE DK ES ER GB GR IT LI LU NI SE N

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

Ι

IE, SI, LT, LV, FI, RO

BR 2000005843 A 20020102 BR 2000-5843 20000703

PRIORITY APPLN. INFO.: EP 1999-112696 A 19990702

OTHER SOURCE(S): MARPAT 134:102214

GI

AΒ Water-sol. fluorescent cyanine dyes, capable of being excited by inexpensive light-emitting diodes or diode lasers and of conjugating with a wide variety of biomols., have the structure I [Q = conjugated connecting group; R1, R2 = H, C1-4 (sulfo)alkyl; R3-R5 = H, SO3H, C1-4 sulfoalkyl, SO2NH(CH2)mW(CH2)nZ; W = direct link, SO2NH, O, CO2, CONH; X1, X2 = O, S, CMe2, C:CH2; Y1, Y2 = benzo, naphtho; Z is or contains a functional group capable of bonding to biomols.; m, n = 0-12; m + n = 1-12] or its salt. Thus, K 2,3,3-trimethyl-3H-indole-5sulfonate was converted with PCl5 and POCl3 to the 5-sulfonyl chloride, which was condensed with glycine tert-Bu ester, and the product was alkylated with 1,4-butane sultone to give 5-[[(carboxymethyl)amino]sulfony 1]-2,3,3-trimethyl-1-(4-sulfobutyl)-3H-indolium inner salt (II). 2,3,3-Trimethyl-5-sulfo-1-(4-sulfobutyl)-3H-indolium inner salt was treated first with PhNHCH:NPh and then with II to give a I [Q = CH:CHCH:, R1 = R2 = (CH2)4SO3H; R3 = 5-SO3H, R4 = R5 = H, W = direct link, X1 = X2 =CMe2, Y1 = Y2 = benzo, Z = CO2H, M = 0, M = 1.

IT 316829-76-2P 316829-77-3P 316829-78-4P 316829-79-5P 316829-80-8P 316829-81-9P 316829-82-0P 316829-83-1P 316829-84-2P 316829-85-3P 316829-86-4P 316829-87-5P 316829-88-6P 316829-89-7P 316829-90-0P 316829-91-1P 316829-92-2P 316829-93-3P

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316829-94-4P 316829-95-5P 316829-96-6P
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316830-07-6P 316830-08-7P 316830-09-8P
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316830-41-8P 316830-42-9P 316830-43-0P
316830-44-1P 316830-45-2P 316830-46-3P
316830-47-4P 316830-48-5P 316830-49-6P
316830-50-9P 316830-51-0P 316830-52-1P
316830-53-2P 316830-54-3P 316830-55-4P
316830-56-5P 316830-57-6P 316830-58-7P
316830-59-8P 316830-60-1P 316830-61-2P
RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
(Analytical study); PREP (Preparation); USES (Uses)
   (prepn. of fluorescent cyanine dye labels contq. a sulfonamido linker
   arm)
316829-76-2 HCAPLUS
3H-Indolium, 2-[3-[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-
dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-
sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)
```

RN

CN

RN 316829-77-3 HCAPLUS
CN 3H-Indolium, 2-[3-[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316829-78-4 HCAPLUS

CN 3H-Indolium, 2-[3-[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316829-79-5 HCAPLUS

CN 3H-Indolium, 2-[3-[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316829-80-8 HCAPLUS

CN 3H-Indolium, 2-[3-[5-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$H_{2}N-(CH_{2})_{4}-NH-S$$
 $H_{2}N-(CH_{2})_{4}-NH-S$ 
 $H_{2}N-(CH_{2})_{4}-NH-S$ 

RN 316829-81-9 HCAPLUS

CN 3H-Indolium, 2-[3-[5-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316829-82-0 HCAPLUS

CN 3H-Indolium, 2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316829-83-1 HCAPLUS

CN 3H-Indolium, 2-[5-[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

~ SO3H

RN 316829-84-2 HCAPLUS

CN 3H-Indolium, 2-[5-[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-

dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316829-85-3 HCAPLUS

3H-Indolium, 2-[5-[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO<sub>2</sub>C- (CH<sub>2</sub>)<sub>3</sub>-NH-S Me Me Me Me Me

PAGE 1-B

-- so3-

SO3H

RN 316829-86-4 HCAPLUS

CN 3H-Indolium, 2-[5-[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-

dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO<sub>3</sub>S- (CH<sub>2</sub>)<sub>4</sub> (CH<sub>2</sub>)<sub>4</sub> (CH<sub>2</sub>)<sub>4</sub> HO<sub>2</sub>C- (CH<sub>2</sub>)<sub>5</sub>-NH-S Me Me Me Me

PAGE 1-B

-- so<sub>3</sub>-

SO3H

RN 316829-87-5 HCAPLUS

CN 3H-Indolium, 2-[5-[5-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- so<sub>3</sub>-

SO3H

RN 316829-88-6 HCAPLUS

CN 3H-Indolium, 2-[5-[5-[{(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$H_{2}N-(CH_{2})_{6}-NH-S$$
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 
 $M_{e}$ 

PAGE 1-B

-- so<sub>3</sub>-

SO<sub>3</sub>H

RN

316829-89-7 HCAPLUS 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-CN indol-2-ylidene]-1,3-pentadienyl]-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A -03S- (CH2)4  $(CH_2)_4 - SO_3H$ CH: = CH— CH== CH— CH= Me Me Ме Me

PAGE 1-B

<u></u> 503H

RN 316829-90-0 HCAPLUS

3H-Indolium, 2-[2-[3-[[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-CNdimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

 $- CH_2 - CO_2H$ 

RN

316829-91-1 HCAPLUS 3H-Indolium, 2-[2-[3-[[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-CNdimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) INDEX NAME)

PAGE 1-B

- CH2- CH2- CO2H

RN 316829-92-2 HCAPLUS

3H-Indolium, 2-[2-[3-[[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-CN 3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

-(CH<sub>2</sub>)<sub>3</sub>-CO<sub>2</sub>H

RN 316829-93-3 HCAPLUS

CN 3H-Indolium, 2-[2-[3-[[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

-(CH<sub>2</sub>)<sub>5</sub>-CO<sub>2</sub>H

RN 316829-94-4 HCAPLUS

CN 3H-Indolium, 2-[2-[3-[[5-{[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

 $(CH_2)_4 - SO_3H$ 

- (CH<sub>2</sub>)<sub>4</sub>-NH<sub>2</sub>

RN 316829-95-5 HCAPLUS

-03S- (CH2)4

CN 3H-Indolium, 2-[2-[3-[[5-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- (CH<sub>2</sub>)<sub>6</sub>- NH<sub>2</sub>

RN 316829-96-6 HCAPLUS

CN 3H-Indolium, 2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- (CH<sub>2</sub>)<sub>6</sub>-OH

RN 316829-97-7 HCAPLUS

CN lH-Benz[e]indolium, 2-[3-[6-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$(CH_2)_4 - SO_3H$$
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 

RN 316829-99-9 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[6-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$(CH_2)_4 - SO_3H$$
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 

RN 316830-00-9 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[6-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-01-0 HCAPLUS

CN lH-Benz[e]indolium, 2-[3-[6-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$(CH_2)_4 - SO_3H$$
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 
 $(CH_2)_4 - SO_3H$ 

RN 316830-02-1 HCAPLUS

CN lH-Benz[e]indolium, 2-[3-[6-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-03-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[6-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$H_2N-$$
 (CH<sub>2</sub>)<sub>4</sub>-SO<sub>3</sub>H (CH<sub>2</sub>)<sub>4</sub>-SO<sub>3</sub>- Me Me Me Me Me SO<sub>3</sub>H

RN 316830-04-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[1,3-dihydro-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1-propenyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO- (CH<sub>2</sub>) 
$$_{6}$$
-NH-S

N

O

HE

CH

CH

CH

CH

CH

CH

CH

N

Me

Me

SO<sub>3</sub>H

RN 316830-05-4 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[((carboxymethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-06-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-07-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-08-7 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-09-8 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

(CH<sub>2</sub>)<sub>4</sub>-SO<sub>3</sub>H

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RN 316830-10-1 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[6-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A 
$$(CH_2)_4 - SO_3H$$

$$H_2N - (CH_2)_6 - NH - S$$

$$O$$

$$O$$

$$N$$

$$CH - CH = CH - CH = CH$$

$$-O_3S - (CH_2)_4$$

PAGE 1-B

RN 316830-11-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[1,3-dihydro-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

HO- (CH<sub>2</sub>) 
$$_{6}$$
- NH-  $_{0}$   $_{0}$   $_{0}$   $_{1}$   $_{1}$   $_{1}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{2}$   $_{3}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{3}$   $_{4}$   $_{2}$   $_{3}$   $_{3}$   $_{4}$   $_{4}$   $_{2}$   $_{3}$   $_{3}$   $_{4}$ 

RN 316830-13-4 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO3S 
$$\sim$$
 CH  $\sim$  CH  $\sim$ 

PAGE 1-B

RN 316830-14-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

HO3S 
$$\frac{+}{N}$$
 CH  $=$  CH  $=$ 

- NH- CH2- CH2- CO2H

RN 316830-15-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

- NH- (CH<sub>2</sub>)<sub>3</sub>- CO<sub>2</sub>H

RN 316830-16-7 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

HO3S 
$$\sim$$
 CH  $\sim$  CH  $\sim$ 

-- NH- (CH<sub>2</sub>)<sub>5</sub>- CO<sub>2</sub>H

RN 316830-17-8 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

HO3S 
$$\frac{\text{CH}_2)_4-\text{SO}_3^-}{\text{Me}}$$
  $\frac{\text{Me}}{\text{Me}}$   $\frac{\text{Me}}{\text{Me}}$   $\frac{\text{Me}}{\text{Ne}}$   $\frac{\text{Me}}$ 

PAGE 1-B

-- NH- (CH<sub>2</sub>)<sub>4</sub>-- NH<sub>2</sub>

RN 316830-18-9 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[6-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-1,1-dimethyl-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

HO3S 
$$\sim$$
 CH  $\sim$  CH  $\sim$ 

-NH-(CH<sub>2</sub>)<sub>6</sub>-NH<sub>2</sub>

RN 316830-19-0 HCAPLUS

CN lH-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-2H-benz[e]indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- NH- (CH<sub>2</sub>)<sub>6</sub>- OH

RN 316830-20-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-21-4 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-22-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-23-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-24-7 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-25-8 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[5-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-26-9 HCAPLUS

CN lH-Benz[e]indolium, 2-[3-[1,3-dihydro-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-27-0 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

SO3H

RN 316830-28-1 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-29-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-30-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

$$HO_3S - (CH_2)_4$$
 $N$ 
 $CH - CH = CH - CH = CH$ 
 $HO_2C - (CH_2)_5 - NH - S$ 
 $Me$ 
 $Me$ 
 $Me$ 
 $Me$ 
 $Me$ 
 $Me$ 
 $Me$ 

PAGE 1-B

RN 316830-31-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-32-7 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[5-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-33-8 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[1,3-dihydro-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

∑SO3H

RN 316830-34-9 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(carboxymethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-35-0 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(2-carboxyethyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-36-1 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(3-carboxypropyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-37-2 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(5-carboxypentyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-38-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(4-aminobutyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-39-4 HCAPLUS

CN 1H-Benz[e]indolium, 2-[2-[3-[[5-[[(6-aminohexyl)amino]sulfonyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, innersalt (9CI) (CA INDEX NAME)

PAGE 1-B

RN 316830-40-7 HCAPLUS

CN lH-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-5-[[(6-hydroxyhexyl)amino]sulfonyl]-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-6-sulfo-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 316830-41-8 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(carboxymethyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-42-9 HCAPLUS

CN 1H-Benz[e]indolium, 6-[((2-carboxyethyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-43-0 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(3-carboxypropyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-44-1 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(5-carboxypentyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-45-2 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(4-aminobutyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO3S Me Me Me 
$$^{\text{CH}_2)}_{4} = \text{SO}_{3}\text{H}$$

HO4S Me Me  $^{\text{H}_2}_{4} = \text{CH}_{2}$ 
 $^{\text{H}_2}_{4} = \text{NH}_{2}$ 
 $^{\text{H}_3}_{5} = \text{NH}_{2}$ 
 $^{\text{H}_2}_{6} = \text{NH}_{2}$ 

RN 316830-46-3 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(6-aminohexyl)amino]sulfonyl]-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-47-4 HCAPLUS

CN 1H-Benz[e]indolium, 2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 316830-48-5 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(carboxymethyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

-- со2н

RN 316830-49-6 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(2-carboxyethyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

— сн<sub>2</sub>— со<sub>2</sub>н

RN 316830-50-9 HCAPLUS

CN 1H-Benz[e]indolium, 6-[((3-carboxypropyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

-(CH<sub>2</sub>)<sub>3</sub>-CO<sub>2</sub>H

RN 316830-51-0 HCAPLUS

CN lH-Benz[e]indolium, 6-[[(5-carboxypentyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

-(CH<sub>2</sub>)<sub>5</sub>-CO<sub>2</sub>H

RN 316830-52-1 HCAPLUS

CN lH-Benz[e]indolium, 6-[((4-aminobutyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

## - (CH<sub>2</sub>)<sub>4</sub> - NH<sub>2</sub>

RN 316830-53-2 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(6-aminohexyl)amino]sulfonyl]-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

## - (CH<sub>2</sub>)<sub>6</sub>- NH<sub>2</sub>

RN 316830-54-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- (CH<sub>2</sub>)<sub>6</sub>-OH

RN 316830-55-4 HCAPLUS

CN lH-Benz[e]indolium, 6-[[(carboxymethyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

(CH<sub>2</sub>)<sub>4</sub>-SO<sub>3</sub>H

Me Me O
S-NH

-O<sub>3</sub>S-(CH<sub>2</sub>)<sub>4</sub>

PAGE 1-B

— cн<sub>2</sub>- co<sub>2</sub>н

RN 316830-56-5 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(2-carboxyethyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

HO3S Me Me 
$$CH-CH$$
 $CH=CH$ 
 $CH$ 
 $CH$ 

-- CH<sub>2</sub>-- CH<sub>2</sub>-- CO<sub>2</sub>H

RN 316830-57-6 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(3-carboxypropyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

- (CH<sub>2</sub>)<sub>3</sub>-CO<sub>2</sub>H

RN 316830-58-7 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(5-carboxypentyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- (CH<sub>2</sub>)<sub>5</sub>-CO<sub>2</sub>H

RN 316830-59-8 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(4-aminobutyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

- (CH<sub>2</sub>)<sub>4</sub> - NH<sub>2</sub>

RN 316830-60-1 HCAPLUS

CN 1H-Benz[e]indolium, 6-[[(6-aminohexyl)amino]sulfonyl]-2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- (CH<sub>2</sub>)<sub>6</sub>-NH<sub>2</sub>

RN 316830-61-2 HCAPLUS

Me

Me

CN 1H-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2-ylidene]ethylidene]-1-cyclohexen-1-yl}ethenyl]-6-[[(6-hydroxyhexyl)amino]sulfonyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

(CH2)  $_4$ -SO3H

Me Me O
S-NH

HO3S

-03S- (CH2) 4

PAGE 1-B

- (CH<sub>2</sub>)<sub>6</sub>-OH

REFERENCE COUNT:

6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

# MAUPIN 09/829,467

### => d ibib abs hitstr 160 8

L60 ANSWER 8 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2000:825711 HCAPLUS

DOCUMENT NUMBER: 134:159683

TITLE: Identification of single fluorescently

labelled mononucleotide molecules in

solution by spectrally resolved time-correlated

single-photon counting

AUTHOR(S): Herten, D. P.; Tinnefeld, P.; Sauer, M.

CORPORATE SOURCE: Physikalisch-Chemisches Institut, Universitat

Heidelberg, Heidelberg, 69120, Germany

SOURCE: Applied Physics B: Lasers and Optics (2000), 71(5),

765-771 CODEN: APBOEM; ISSN: 0946-2171

Springer-Verlag

PUBLISHER: Springe:
DOCUMENT TYPE: Journal
LANGUAGE: English

AB We describe a method to identify single dye-labeled mononucleotide mols. in soln. with high classification probability based on confocal microscopy in combination with spectrally and time-resolved fluorescence detection with two detectors. For efficient excitation of the labeled mononucleotide mols. JA133-dUTP, JA169-dUTP, Cy5-dCTP, and JA242-dUTP a short-pulse diode laser emitting at 634 nm with a repetition rate of 64 MHz was applied. The time-resolved fluorescence signals of individual mols. were analyzed and identified by a max. likelihood estimator (MLE). Scatter plots of spectrally and time-resolved fluorescence data demonstrated the existence of four distinct populations with sym. shape. The distributions of each of the mononucleotide conjugates were detd. by fitting a superposition of two independent Gaussians. Taking only those single-mol. bursts which contain more than 50 photon counts, three labeled mononucleotide mols. were identified in soln. by spectrally and time-resolved fluorescence spectroscopy with a probability of correct classification of .apprxeq. 99%.

IT 325747-77-1

RL: ANT (Analyte); BSU (Biological study, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study)

(identification of single **fluorescently** labeled **mononucleotide** mols. in soln. by spectrally resolved

time-correlated single-photon counting)

RN 325747-77-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[[3-[4-amino-1-[2-deoxy-5-O-[hydroxy([hydroxy(phosphonooxy)phosphinyl]oxy]phosphinyl]-.beta.-D-erythro-pentofuranosyl]-1,2-dihydro-2-oxo-5-pyrimidinyl]-2-propynyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

H2N\_

PAGE 1-B

REFERENCE COUNT:

23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

# MAUPIN 09/829,467

=> d ibib abs hitstr 160 9

L60 ANSWER 9 OF 17 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2000:790577 HCAPLUS

DOCUMENT NUMBER: 133:351506

TITLE: Aza-benzazolium-containing cyanine dyes and their use

in fluorescent biological stains

INVENTOR(S): Haugland, Richard P.; Yue, Stephen T.

PATENT ASSIGNEE(S): Molecular Probes, Inc., USA

SOURCE: PCT Int. Appl., 87 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2000066664 A1 20001109 WO 2000-US11549 20000426

W: AU, CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

PT, SE PRIORITY APPLN. INFO.:

US 1999-131782P P 19990430 US 1999-158859P P 19991012

OTHER SOURCE(S): CASREACT 133:351506

AB Unsym. cyanine dyes that incorporate an aza-benzazolium ring moiety are disclosed, including cyanine dyes substituted by a cationic side chain, monomeric and dimeric cyanine dyes, chem. reactive cyanine dyes, and conjugates of cyanine dyes. The dyes are virtually non-fluorescent when dild. in aq. soln., but exhibit bright fluorescence when assocd. with nucleic acid polymers such as DNA or RNA, or when assocd. with detergent-complexed proteins. A variety of applications are described for detection and quantitation of nucleic acids and detergent-complexed proteins in a variety of samples, including solns., electrophoretic gels, cells, and microorganisms.

IT 305802-24-8P

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); IMF (Industrial manufacture); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)

(dye; prodn. of azabenzazolium cyanine dyes for fluorescent biol. stains)

RN 305802-24-8 HCAPLUS

CN Quinolinium, 1-methyl-4-[3-(3-methyl-2(3H)-benzothiazolylidene)-2-phenyl-1-propenyl]-, salt with 4-methylbenzenesulfonic acid (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 305802-23-7 CMF C27 H23 N2 S

CM 2

CRN 16722-51-3 CMF C7 H7 O3 S

REFERENCE COUNT:

8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

HCAPLUS COPYRIGHT 2002 ACS L60 ANSWER 10 OF 17

ACCESSION NUMBER:

1999:794372 HCAPLUS

DOCUMENT NUMBER:

132:35989

TITLE:

Preparation of cyanine dye activating group

with improved coupling selectivity to label chain

terminators in nucleotide sequencing Shen, Gene G.-Y.; Dobashi, Thomas S.

INVENTOR(S): PATENT ASSIGNEE(S):

Beckman Instruments, Inc., USA

SOURCE:

U.S., 19 pp.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ----------------US 6002003 19991214 US 1998-59900 19980414 Α

OTHER SOURCE(S):

MARPAT 132:35989

GI

AΒ Activating groups for cyanine dyes I (R1 = R2 = H; R1 = phthalimido; R2 = R3; X = CH:CHCH:CHCH:, CH:CHCH:CHCH:Ohch:CHCH:) used to label chain terminators in nucleotide sequencing, based on N-hydroxyphthalimide, are disclosed. From these activating groups, activated dyes of the present invention are prepd. which react with the derivatized nucleotide chain terminators to give a labeled chain terminator of the present Invention. The activating groups of the present Invention allow the dye-chain terminator reaction to occur at a much higher yield and with much greater selectivity for the mono-substituted product, compared with the prior art.

#### ΙT 252255-44-0P 252255-45-1P 252255-48-4P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. of cyanine dye activating group with improved

coupling selectivity to label chain terminators in nucleotide sequencing)

RN

252255-44-0 HCAPLUS 3H-Indolium, 1-(5-carboxypentyl)-2-[5-[1,3-dihydro-3,3-dimethyl-1-[6-oxo-6-CN [[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2furanyl]-5-pyrimidinyl]-2-propynyl]amino]hexyl]-5-sulfo-2H-indol-2ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, inner salt, monopotassium trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

K

3 Na

#### RN 252255-45-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-1-[6-oxo-6-[[3-[1,2,3,4tetrahydro-2, 4-dioxo-1-[(2R,5S)-tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]hexyl]-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3dimethyl-1-[6-oxo-6-[3-[1,2,3,4-tetrahydro-2,4-dioxo-1-[(2R,5S)tetrahydro-5-(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]hexyl]-5sulfo-, inner salt, monopotassium hexasodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 1-A

K

PAGE 1-C

●6 Na

RN 252255-48-4 HCAPLUS CN 3H-Indolium, 1-[6-[[3-[4-amino-1,2-dihydro-2-oxo-1-[(2R,5S)-tetrahydro-5(3,5,7,7-tetrahydroxy-3,5,7-trioxido-2,4,6-trioxa-3,5,7-triphosphahept-1-yl)-2-furanyl]-5-pyrimidinyl]-2-propynyl]amino]-6-oxohexyl]-2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, inner salt, monopotassium trisodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

K

PAGE 1-B

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L60 ANSWER 11 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:708965 HCAPLUS

DOCUMENT NUMBER: 129:335785

TITLE: Acid-labile and enzymically cleavable dye

conjugates for diagnosis with near-IR

radiation and for therapy

INVENTOR(S):
Licha, Kai; Riefke, Bjoern; Semmler, Wolfhard;

Wrasidlo, Wolfgang

PATENT ASSIGNEE(S): Institut fuer Diagnostikforschung G.m.b.H. an der

Freien Universitaet Berlin, Germany

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	rent	NO.		KI	DN	DATE				AP	PLI	CATI	ON 1	10.	DATE			
	9847 9847									WO	19	98-D	E100	)1	1998	0402		
						JP,												
	RW:	AT, PT,		CH,	CY,	DE,	DK,	ES,	FI	,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,
DE	1971	7904		A:	1	1998	1029			DE	19	97-1	9717	7904	1997	0423		
AU	9879	057		A:	1	1998	1113			ΑU	19	98-7	9057	7	1998	0402		
AU	7337	57		B	2	2001	0524											
EP	9880	60		A2	2	2000	0329			ΕP	19	98-9	2921	L2	1998	0402		
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GE	3, (	GR,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,
		IE,																
JP	2001	52153	30	T	2	2001	1106			JP	19	98-5	4471	L5	1998	0402		
NO	9905	181		Α		1999	1022			NO	19	99-5	181		1999	1022		
PRIORITY	Y APP	LN.	INFO	. :					DΕ	19	97-	1971	7904	1 A	1997	0423		
									WO	19	98-	DE10	01	W	1998	0402		
OTHER SO	OURCE	(S):			MAR	PAT	129:3	3357	85									

AB Dyes which fluoresce in the near-IR spectral region are provided, the fluorescence of which is quenched by coupling via a cleavable linker to arom. compds. (e.g. dyes, drugs), antibodies, antibody fragments, or other proteins. Cleavage of such a construct in vivo at a target site (e.g. a tumor or focus of inflammation) leads to an increase in near-IR fluorescence, which can be detected even at deep sites owing to the high

# MAUPIN 09/829,467

transparency of tissues to near-IR radiation. Suitable dyes include tetrapyrrole, tetraazapyrrole, xanthine, phenoxazine, phenothiazine, and esp. polymethine dyes such as cyanine dyes. Drug-dye conjugates in which the therapeutic activity of the drug is masked by coupling to the dye may serve as prodrugs which, after administration, are cleaved at a target site to release the active agent, as well as the fluorescent dye which may act as photosensitizer, at the site. The linker may be acid labile, i.e. cleavable at the low pH characteristic of tumors and sites of bacterial inflammation, or cleavable by enzymes which occur in diseased tissues, e.g. bacterial enzymes. Thus, a cyanine dye, 5-(1-oxoethyl)-1,1'-(4-sulfobutyl)indotricarbocyanine Na salt (I) was prepd. by reaction of 4-hydrazinophenyl Me ketone with 3-methyl-2-butanone followed by 1,4-butanesultone to form 5-(1-oxoethyl)-1-(4-sulfobutyl)-2,3,3-trimethyl-3H-indolenine and further reaction of this compd. with glutaconaldehyde dianil-HCl. Reaction of I with 4carboxyphenylsulfonylhydrazine followed by N-hydroxysuccinimide and DCCD produced an acid-labile N-hydroxysuccinimidyl ester, which was coupled to anti-melanoma monoclonal antibody 9.2.27; the antibody conjugate had a fluorescence quantum yield of 0.1%.

215114-68-4P 215114-71-9P 215114-72-0P ΙT

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(acid-labile and enzymically cleavable dye conjugates for diagnosis with near-IR radiation and for therapy)

RN

215114-68-4 HCAPLUS
3H-Indolium, 5-[1-[[(4-carboxyphenyl)sulfonyl]hydrazono]ethyl]-2-[7-[1,3-CN dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

215114-71-9 HCAPLUS RN

3H-Indolium, 5-[1-[[[4-[[(2,5-dioxo-1-pyrrolidinyl)oxy]carbonyl]phenyl]sul CN

fonyl]hydrazono]ethyl]-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

● Na

PAGE 1-B

RN 215114-72-0 HCAPLUS

CN 3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-5-[1-[[(4-isothiocyanatophenyl)sulfonyl]hydrazono]ethyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

L60 ANSWER 12 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1998:604701 HCAPLUS

DOCUMENT NUMBER:

129:216857

TITLE:

Preparation of indocarbocyanine and benzindocarbocyanine dye-linked

phosphoramidites

INVENTOR(S):

Brush, Charles K.; Anderson, Eric Dean

PATENT ASSIGNEE(S):

Pharmacia Biotech Inc., USA

SOURCE:

U.S., 23 pp., Cont.-in-part of U.S. Ser. No. 712,505,

abandoned. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE		APPLICATION NO.	DATE
US 5808044	A	19980915		US 1997-799593	19970210
US 5556959	Α	19960917		US 1994-265569	19940624
PRIORITY APPLN. INFO.	:		US	1993-7444	19930122
			US	1994-265569	19940624
			US	1996-712505	19960911

OTHER SOURCE(S):

GΙ

MARPAT 129:216857

Me Me 
$$(CH=CH)_p-CH$$
 Me Me  $(CR=CH)_p-CH$  Me Me  $(CR=CH)_p-CH$  Me

AB The title compds. [I; R = H, trityl,4,4'-dimethoxytrityl, or acyl groups group or is an H; Ra = a phosphorami

alkyl; R4, R5 = H, lower alkyl, acyl, —en-enen-en-or (enzyp1CO2(CH2)qMe; where p1, q = an integer from 0 to 4; m, n = an integer from 0 to 10; p = an1, 2, or 3; X- = a neg. ion], which are useful for fluorescent, non-radioactive labeling of oligonucleotides in automated DNA sequencing, in situ detection of hybridization, etc., are prepd. Oligonucleotides I (R = aryl group-contg. moiety, which does not interfere with the attachment of an oligonucleotides at the Ra position; Ra = an oligonucleotide; R1a, R1b, R4, R5, m, n, p, X- = same as above) and method for linking a fluorescent label to an oligonucleotide are also claimed. Thus, a soln. of 28.15 g 1,3,3-trimethoxypropene in MeCN was added dropwise at 100.degree. over 1 h to a soln. of 110 g 1-(1'-acetoxypropy1)-2,3,3-trimethyl-(3H)-indolinium iodide, 24 mL Et3N, and 8 mL AcOH in MeCN, refluxed for 2 h, cooled, and evapd. to a gum,

protecting

### MAUPIN 09/829,467

which was dissolved in 2 M aq. HCl in 50:50 H2O/MeOH and stirred at room temp. overnight to give, after workup and silica gel chromatog., 60 g 1,1'-bis(1''-hydroxy-3''-propyl)-3,3,3',3'-tetramethylindodicarbocyan ine chloride I (p = 2, n = m = 1, R = Ra = R1a = R1b = R4 = R5 = H).This compd. (6 g) was tritylated by monomethoxytrityl chloride in pyridine to give 3-4 g I (r = 2, n = m = 1, R = monomethoxytrityl, Ra = Rla = Rlb = RR4 = R5 = H), which (4.0 g) was condensed with 2.32 g NCCH2CH2P[N(CHMe2)2]2 in MeCN to give the title compd. I (p = 2, n = m = 1, R = monomethoxytrityl, Ra = P[N(CHMe2)2]2CH2CH2CN, Rla = Rlb = R4 = R5 = H).

182873-67-2P 212563-53-6P 212563-54-7P IT 212563-58-1P

> RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of indocarbocyanine and benzindocarbocyanine dye-linked phosphoramidites fluorescent, non-radioactive labeling of oligonucleotides)

RN

182873-67-2 HCAPLUS 3H-Indolium, 2-[5-[1-[3-[[[bis(1-methylethyl)amino](2-CN cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1,3-pentadienyl]-1-[3-[(4-methoxyphenyl)diphenylmethoxy]propyl]-3,3-dimethyl-, chloride (9CI) (CA INDEX NAME)

● cl-

RN 212563-53-6 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-[3-[[[bis(1-methylethyl)amino](2cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-[3-[(4-methoxyphenyl)diphenylmethoxy]propyl]-1,1-dimethyl-, chloride (9CI) (CA INDEX NAME)

● cl-

RN 212563-54-7 HCAPLUS

CN 1H-Benz[e]indolium, 3-[3-(acetyloxy)propyl]-2-[5-[3-[3-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-, chloride (9CI) (CA INDEX NAME)

● C1-

RN 212563-58-1 HCAPLUS

CN 1H-Benz[e]indolium, 3-[3-(acetyloxy)propyl]-2-[3-[3-[3-[3-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1-propenyl]-1,1-dimethyl-, chloride (9CI) (CA INDEX NAME)

● c1-

IT 182873-76-3P 182873-82-1P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of indocarbocyanine and benzindocarbocyanine dye-linked phosphoramidites fluorescent, non-radioactive labeling of oligonucleotides)

RN 182873-76-3 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[3-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1-[3-[(4-methoxyphenyl)diphenylmethoxy]propyl]-3,3-dimethyl-, chloride (9CI) (CA INDEX NAME)

• c1-

RN 182873-82-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[3-[[[bis(1-methylethyl)amino](2-cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-[3-(2,2-dimethyl-1-oxopropoxy)propyl]-3,3-

dimethyl-, chloride (9CI) (CA INDEX NAME)

PAGE 1-A

NC-CH<sub>2</sub>-CH<sub>2</sub>-O-P-O-(CH<sub>2</sub>)<sub>3</sub>

CH-CH=CH-CH=CH-CH
Me
Me
Me
Me

• c1-

PAGE 1-B

— Bu−t

L60 ANSWER 13 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:466230 HCAPLUS

DOCUMENT NUMBER: 127:172704

TITLE: Stability, specificity and fluorescence brightness of

multiply-labeled fluorescent DNA

probes

AUTHOR(S): Randolph, John B.; Waggoner, Alan S.

CORPORATE SOURCE: Center Light Microscope Imaging Biotechnology

Department Chemistry, Carnegie Mellon University,

Pittsburgh, PA, 15213, USA

SOURCE: Nucleic Acids Research (1997), 25(14), 2923-2929

CODEN: NARHAD; ISSN: 0305-1048

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal LANGUAGE: English

AB In this work, we studied the fluorescence and hybridization of multiply-labeled DNA probes which have the hydrophilic fluorophore

1-(.epsilon.-carboxypentynyl)-1'-ethyl-3,3,3',3'tetramethylindocarbocyanine-5,5'-disulfonate (Cy3)

attached via either a short or long linker at the C-5 position of deoxyuridine. We describe the effects of labeling d., fluorophore charge and linker length upon five properties of the probe: fluorescence intensity, the change in fluorescence upon duplex formation, the quantum yield of fluorescence (.PHI.f), probe-target stability and specificity. For the hydrophilic dye Cy3, we have demonstrated that the fluorescence intensity and .PHI.f are maximized when labeling every 6th base using the long linker. With a less hydrophilic dye, a labeling d. this high could not be achieved without serious quenching of the fluorescence. The target specificity of multiply-labeled DNA probes was just as high as compared to the unmodified control probe, however, a less stable probe-target duplex is formed that exhibits a lower melting temp. A mechanism that accounts for this destabilization is proposed which is consistent with our data. It involves dye-dye and dye-nucleotide interactions which appear to stabilize a single-stranded conformation of the probe.

IT 194159-49-4 194159-50-7

RL: ARU (Analytical role, unclassified); ANST (Analytical study) (stability, specificity, and fluorescence brightness of multiply-labeled **fluorescent DNA** probes)

RN 194159-49-4 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[5-[[6-[[3-[1-(2-deoxy-3-O-phosphono-.beta.-D-erythro-pentofuranosyl)-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-1-oxo-2-propenyl]amino]hexyl]amino]pentyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

# PAGE 1-B

RN 194159-50-7 HCAPLUS

3H-Indolium, 2-[3-[1-[5-[[2-[[3-[1-(2-deoxy-3-O-phosphono-.beta.-D-erythro-pentofuranosyl)-1,2,3,4-tetrahydro-2,4-dioxo-5-pyrimidinyl]-1-oxo-2-propenyl]amino]ethyl]amino]pentyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.

L60 ANSWER 14 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:610223 HCAPLUS

DOCUMENT NUMBER: 125:301494

TITLE: Preparation of indocarbocyanine dye-linked

phosphoramidites

INVENTOR(S): Brush, Charles K.; Anderson, Eric D. PATENT ASSIGNEE(S): Pharmacia P-L Biochemicals Inc., USA

SOURCE: U.S., 15 pp., Cont. of U.S. Ser. No. 7,444, abandoned.

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	<del></del>			
US 5556959	A	19960917	US 1994-265569	19940624
US 5808044	A	19980915	US 1997-799593	19970210
PRIORITY APPLN. IN	IFO.:		US 1993-7444	19930122
			US 1994-265569	19940624
			US 1996-712505	19960911

OTHER SOURCE(S): MARPAT 125:301494

GI

Me Me Me Me Me Me 
$$R^{5}$$
 $R^{5}$ 
 $R^{5}$ 
 $R^{6}$ 
 $R^{7}$ 
 $R$ 

AB The title compds. [I; R = H, trityl, 4-monomethoxytrityl, 4,4'-dimethoxytrityl, or acyl groups and R may be used as a protecting group or is an H; Ra = a phosphoramidite; Rla, Rlb = H, lower alkyl; R4, R5 = H, lower alkyl, acyl, or (CH2)pCO2(CH2)qMe; where p, q = an integer from 0 to 4; m, n = an integer from 0 to 10; r = 1, 2, or 3; X-= a neg. ion], which are useful for fluorescent, non-radioactive labeling of oligonucleotides in automated DNA sequencing, in situ detection of hybridization, etc., are prepd. Oligonucleotides I (R = aryl group-contg. moiety, which does not interfere with the attachment of an oligonucleotides at the Ra position; Ra = an oligonucleotide; Rla, Rlb, R4, R5, m, n, r, X-= same as above) and method for linking a fluorescent label to an oligonucleotide are also claimed. Thus, a soln. of 28.15 g 1,3,3-trimethoxypropene in MeCN was added dropwise at 100.degree. over 1 h to a soln. of 110 g 1-(1'-acetoxypropyl)-2,3,3-trimethyl-(3H)-indolinium iodide, 24 mL Et3N, and 8 mL AcOH in MeCN, refluxed for 2 h, cooled, and evapd. to a qum, which was dissolved in 2 M aq. HCl in 50:50 H2O/MeOH and stirred at room temp. overnight to give, after workup and silica gel chromatog., 60

Ι

g1,1'-bis(1''-hydroxy-3''-propyl)-3,3,3',3'tetramethylindodicarbocyanine chloride I (r = 2, n = m = 1, R = Ra =  $R1a = R\overline{1}b = R4 = R5 = H$ ). This compd. (6 g) was tritylated by monomethoxytrityl chloride in pyridine to give 3-4 g I (r = 2, n = m = 1, R = monomethoxytrityl, Ra = Rla = Rlb = R4 = R5 = H), which (4.0 g) was condensed with 2.32 g NCCH2CH2P[N(CHMe2)2]2 in MeCN to give the title compd. I (r = 2, n = m = 1, R = monomethoxytrityl, Ra = monomethoxytrityl, RP[N(CHMe2)2]2CH2CH2CN, R1a = R1b = R4 = R5 = H).182873-67-2P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of indocarbocyanine dye-linked phosphoramidites fluorescent, non-radioactive labeling of oligonucleotides)

RN

ΙT

182873-67-2 HCAPLUS 3H-Indolium, 2-[5-[1-[3-[[[bis(1-methylethyl)amino](2-CN cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1,3-pentadienyl]-1-[3-[(4-methoxyphenyl)diphenylmethoxy]propyl]-3,3-dimethyl-, chloride (9CI) (CA INDEX NAME)

Cl-

#### ΙT 182873-76-3P 182873-82-1P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of indocarbocyanine dye-linked phosphoramidites fluorescent, non-radioactive labeling of oligonucleotides)

RN 182873-76-3 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[3-[[bis(1-methylethyl)amino](2cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2ylidene]-1-propenyl]-1-[3-[(4-methoxyphenyl)diphenylmethoxy]propyl]-3,3dimethyl-, chloride (9CI) (CA INDEX NAME)

• c1-

RN

182873-82-1 HCAPLUS 3H-Indolium, 2-[5-[1-[3-[[[bis(1-methylethyl)amino](2-CN cyanoethoxy)phosphino]oxy]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-[3-(2,2-dimethyl-1-oxopropoxy)propyl]-3,3dimethyl-, chloride (9CI) (CA INDEX NAME)

● Cl-

PAGE 1-B

— Bu−t

L60 ANSWER 15 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1995:3281 HCAPLUS

DOCUMENT NUMBER: 122:155746

TITLE: Fluorescent dyes for labeling biosubstances for

analysis

INVENTOR(S): Katayose, Mitsuo; Tai, Seiji; Watanabe, Hiroo

PATENT ASSIGNEE(S): Hitachi Chemical Co Ltd, Japan SOURCE: Jpn. Kokai Tokkyo Koho, 8 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

JP 05287209 A2 19931102 JP 1992-88743 19920409

OTHER SOURCE(S): MARPAT 122:155746

GI For diagram(s), see printed CA Issue.

AB The fluorescent dyes I (A1, A2 = benzene, naphthalene, etc.; R1-3 = H, alkyl, alkoxy; X1 = S, O, etc.; X2 = S, O, CO, etc.; and L = polymethylene) and biosubstance (e.g. vitamin, nucleotide, or protein) labeled with the fluorescent dye are prepd. and used for antigen, pharmaceutical, or DNA anal. Four such fluorescent dyes, their sulfonylchloride derivs. and p-aminobenzoic acid adducts, and 3-(4-aminobutyl)morphine labeled with these dyes were prepd. for morphine anal. A labeled DNA probe, i.e. GTTTCCCAGTCACGAC, was also prepd. for DNA sequence anal. The detection of the dye is not affected by heme in blood and is therefore a good test reagent for clin. anal.

IT 154187-63-0P 154187-64-1P 161066-73-5P 161066-74-6P 161098-14-2P 161098-15-3P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of, as label, for biosubstance detn.)

RN 154187-63-0 HCAPLUS

CN Quinolinium, 4-[5-[1-[3-(chlorosulfonyl)propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-, chloride (9CI) (CA INDEX NAME)

RN 154187-64-1 HCAPLUS

CN Quinolinium, 4-[5-[1-[3-[[(4-carboxyphenyl)amino]sulfonyl]propyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-, chloride (9CI) (CA INDEX NAME)

RN 161066-73-5 HCAPLUS

CN 3H-Indolium, 1-[3-(chlorosulfonyl)propyl]-3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-, chloride (9CI) (CA INDEX NAME)

D1-Me

• c1-

RN 161066-74-6 HCAPLUS

CN 3H-Indolium, 1-[3-[((4-carboxyphenyl)amino]sulfonyl]propyl]-3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-, chloride (9CI) (CA INDEX NAME)

D1-Me

RN 161098-14-2 HCAPLUS

CN 3H-Benz[g]indolium, (chlorosulfonyl)-1-[4-(chlorosulfonyl)butyl]-3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzothiopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-, chloride (9CI) (CA INDEX NAME)

● C1-

D1-Me

RN 161098-15-3 HCAPLUS

CN 3H-Benz[g]indolium, 1-[4-[[(4-carboxyphenyl)amino]sulfonyl]butyl](chlorosu lfonyl)-3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzothiopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-, chloride (9CI) (CA INDEX NAME)

# PAGE 2-A

D1-Me

# ● Cl-

IT 158019-49-9P 158019-50-2P 158019-51-3P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of, for morphine detn.)

RN 158019-49-9 HCAPLUS

CN 3H-Indolium, 1-[3-[[[4-[[(5.alpha.,6.alpha.)-7,8-didehydro-4,5-epoxy-6-hydroxy-17-methylmorphinan-3-yl]butyl]amino]carbonyl]phenyl]amino]sulfonyl]propyl]-2-[5-(1-ethyl-4(1H)-quinolinylidene)-1,3-pentadienyl]-3,3-dimethyl-, chloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

Me Me 
$$(CH_2)_3$$
  $H$   $N$   $O$   $O$   $H$   $N$   $O$   $O$ 

● C1-

PAGE 1-B

$$(CH_2)_4$$
  $(CH_2)_4$   $(CH_2)_4$ 

RN 158019-50-2 HCAPLUS

CN Benzothiazolium, 3-[3-[[4-[[4-[(5.alpha.,6.alpha.)-7,8-didehydro-4,5-epoxy-6-hydroxy-17-methylmorphinan-3-yl]butyl]amino]carbonyl]phenyl]amino]sulfonyl]propyl]-2-[5-(1-ethyl-4(1H)-quinolinylidene)-1,3-pentadienyl]-,chloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

● C1-

PAGE 1-B

$$(CH_2)_4$$
 OH OH

RN 158019-51-3 HCAPLUS

CN 3H-Indolium, 1-[3-[[[4-[[[4-[(5.alpha.,6.alpha.),7,8-didehydro-4,5-epoxy-6-hydroxy-17-methylmorphinan-3-yl]butyl]amino]carbonyl]phenyl]amino]sulfonyl]propyl]-3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-, chloride (9CI) (CA INDEX NAME)

# PAGE 2-A

D1-Me

● c1-

# MAUPIN 09/829,467

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L60 ANSWER 16 OF 17 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1994:527040 HCAPLUS

DOCUMENT NUMBER: 121:127040

TITLE: Chromogenic mononucleotide analogs for

DNA probe

INVENTOR(S): Tomita, Yoshinori; Mihara, Cheko; Okamoto, Hisashi

PATENT ASSIGNEE(S): Canon Kk, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 13 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 06122696	A2	19940506	JP 1992-272914	19921012

AB The chromogenic mononucleotide analogs comprise dye, phosphate group, and carbohydrate moiety. Thus, 1-{1,1-dimethyl-2-[7-(1,1-dimethyl-3-ethyl-indolin-2-ylidene)-1,3,5-heptatrienyl]-1H-indolium iodide}-3'-(.beta.-cyanoethyl-N,N-diisopropylphosphoramide)-5'-dimethoxytrityl-2'-deoxy-D-ribose was synthesized. from 2,3,3-trimethyl-indolenin, Et iodide, 1-iodo-5-dimethoxytrityl-2-deoxy-D-ribose, etc., and used for prepg. dye-label DNA probe.

IT 157132-15-5P 157132-16-6P 157132-21-3P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and reaction of, in prepg. chromogenic mononucleotide

analogs for **DNA** probe)

RN 157132-15-5 HCAPLUS

CN 3H-Indolium, 2-[6-[1-[3-O-acetyl-5-O-[bis(4-methoxyphenyl)phenylmethyl]-2-deoxy-.beta.-D-erythro-pentofuranosyl]-2,3-dihydro-3,3-dimethyl-1H-indol-2-yl]-1,3,5-hexatrienyl]-1-ethyl-3,3-dimethyl-, iodide (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 2-A

• I-

RN 157132-16-6 HCAPLUS

CN 3H-Indolium, 2-[6-[1-[5-0-[bis(4-methoxyphenyl)phenylmethyl]-2-deoxy-beta.-D-erythro-pentofuranosyl]-2,3-dihydro-3,3-dimethyl-1H-indol-2-yl]-1,3,5-hexatrienyl]-1-ethyl-3,3-dimethyl-, iodide (9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

PAGE 2-A

• I-

RN 157132-21-3 HCAPLUS

CN 1H-Indole, 2-[7-[1-(2-deoxy-.beta.-D-erythro-pentofuranosyl)-3-methyl-1H-indol-2-yl]-2,4,6-heptatrienylidene]-1-ethyl-2,3-dihydro-3,3-dimethyl-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Double bond geometry unknown.

IT 157132-20-2P

RN 157132-20-2 HCAPLUS

CN 1H-Benz[e]indole, 2-[7-[3-[5-O-[bis(4-methoxyphenyl)phenylmethyl]-2-deoxy-.beta.-D-erythro-pentofuranosyl]-1-methyl-1H-benz[e]indol-2-yl]-2,4,6-heptatrienylidene]-3-ethyl-2,3-dihydro-1,1-dimethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

HCAPLUS COPYRIGHT 2002 ACS L60 ANSWER 17 OF 17 1982:451808 HCAPLUS

ACCESSION NUMBER:

DOCUMENT NUMBER: 97:51808

TITLE: Photographic microquantitation of enzymes

Fuji Photo Film Co., Ltd., Japan PATENT ASSIGNEE(S): SOURCE: Jpn. Kokai Tokkyo Koho, 31 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 57047493	A2	19820318	JP 1980-120600	19800902
JP 61001118	B4	19860114		
EP 48834	A1	19820407	EP 1981-106826	19810901
EP 48834	B1	19850619		
R: CH, DE,	FR, GB			
US 4414325	Α	19831108	US 1981-298814	19810902
PRIORITY APPLN. INFO	. :		JP 1980-120600	19800902
GI				

A synthetic substrate (having a mol. structure that specifically reacts AB with the enzyme to be detd. and also that has a spectral sensitizing mol. structure) is contacted with the enzyme to be detd. Then, either the enzymic reaction product or the unreacted synthetic substrate remaining is reacted with an Ag halide, exposed to the spectrum of light that corresponds to the spectral sensitivity of the substrate, photog. developed, and the concn. of the Ag image and (or) the color developed is detd. as an enzyme activity and (or) the enzymic content of the sample. This method is suitable for detg. protein-decompg. enzymes, peptide-decompg. enzymes, nucleic acid-decompg. enzymes, sugar-decompg. enzymes, and lipid-decompg. enzymes. Thus, 1 mL each of I-modified glycylphenylalaninamide (1 mg/mL) in 0.05M Tris-HCl buffer (pH 8.5) contg. 1% surfactant and bovine pancreas .alpha.-chymotrypsin at 2, 20, and 200 pg/mL in 0.05M Tris-HCl buffer (pH 8.5) were mixed, incubated at 40.degree. for 5 min, and mixed with 0.1 mg tosylamidophenylalanylchloromethylketone to stop the enzyme reaction. Each reaction mixt. was passed through CM-Sephadex C-50, the column washed with 1 mL 0.05M Tris-HCl buffer (pH 8.5) and the eluent and washings collected. The collected liq. (25 .mu.L each) was applied to the unexposed AgBrCl film in a spot 5 mm diam. The film was allowed to stand at room temp. in the dark for 20 min and exposed to a light through Fuji Film Filter SC-66 at 108 lx for 10-3 s, conventionally developed, and the intensity of darkness of the spot detd. The darkness of the spots was

Ι

directly proportional to the concn. of .alpha.-chymotrypsin.

- IT 82458-83-1 82458-91-1
  - RL: BIOL (Biological study)
    - (spectral sensitizing pigment, for enzyme assay)
- RN 82458-83-1 HCAPLUS
- CN Benzothiazolium, 2-[5-[6-carboxy-3-[(tetrahydro-2-furanyl)methyl]-2(3H)-benzothiazolylidene]-3-ethyl-1,3-pentadienyl]-5-chloro-3-[(tetrahydro-2-furanyl)methyl]-, iodide (9CI) (CA INDEX NAME)

• I-

- RN 82458-91-1 HCAPLUS
- CN 1-Imidazolidinepropanoic acid, 4-[4-[5,6-dimethyl-3-[(tetrahydro-2-furanyl)methyl]-2(3H)-benzothiazolylidene]-2-butenylidene]-3-ethyl-5-oxo-2-thioxo-(9CI) (CA INDEX NAME)

Me 
$$\sim$$
 CH2  $\sim$  CH  $\sim$  CH  $\sim$  CH  $\sim$  CH2  $\sim$  C

Generic sends

## MAUPIN 09/829,467

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=> d que 168
L15
         16673 SEA FILE=HCAPLUS ABB=ON PLU=ON ?STYRYL?
         863910 SEA FILE=HCAPLUS ABB=ON PLU=ON (DNA OR ?NUCLEOTID? OR
L16
                NUCLEIC)
L18
          18654 SEA FILE=HCAPLUS ABB=ON PLU=ON (SULFONATES/CT OR SULFONATION/
                CT OR "SULFONIC ACIDS"/CT)
L19
           6154 SEA FILE=HCAPLUS ABB=ON PLU=ON "PHOSPHATES, BIOLOGICAL
               STUDIES"/CT
L34
                STR
                13
               √G2√Cy
14
                            Me√ C√√Me
                            10 @11 12
```

VAR G1=O/S/11
REP G2=(2-9) C
NODE ATTRIBUTES:
CONNECT IS E3 RC AT 8
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 14

STEREO ATTRIBUTES: NONE

SIEKEU	ATTRIBUTE	50: I	NONE		
L36	45490	SEA	FILE=REGISTRY SSS FU	L L34	
L39	4949	SEA	FILE=REGISTRY ABB=ON	PLU=ON	L36 AND ?SULFON?/CNS
L40	639	SEA	FILE=REGISTRY ABB=ON	PLU=ON	L36 AND ?PHOSPH?/CNS
L41	39918	SEA	FILE=REGISTRY ABB=ON	PLU=ON	L36 NOT (L39 OR L40)
L42	561	SEA	FILE=REGISTRY ABB=ON	PLU=ON	L41 AND OC4/ES
L46	8330	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L16(3A) FLUORES?
L50	68099	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L15 OR ?CYANINE?
L61	39357	SEA	FILE=REGISTRY ABB=ON	PLU=ON	L41 NOT L42
L62	15024	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L61
L63	266	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L46 AND L62
L64	74	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L63 AND ?CONJUGAT?
L65	21	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L64 AND L50
L66	26	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L64 AND ((L18 OR L19) OR
		?SUI	LFON? OR ?PHOSPH?)		
L67	8	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L65 AND L66
L68	18	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L66 NOT L67

# MAUPIN 09/829,467

=> d ibib abs hitstr 168 1

L68 ANSWER 1 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:172235 HCAPLUS

DOCUMENT NUMBER: 136:213182

TITLE: Methods employing fluorescence quenching by metal

surfaces

INVENTOR(S): Dubertret, Benoit; Calame, Michel; Libchaber, Albert

PATENT ASSIGNEE(S): The Rockefeller University, USA

SOURCE: PCT Int. Appl., 62 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2002018951 A2 20020307 WO 2001-US41941 20010829

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO:

US 2000-228728P P 20000829

US 2001-280350P P 20010330
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AΒ The invention is broadly related to methods for sensitively detecting proximity changes in systems that utilize an interacting fluorophore and quencher. In such methods, a metal surface is used as the quencher. metal surface may be a particle or film, such as nanoparticles or a coating, resp. Such systems provide an increase in sensitivity over previously-described quenchers, offering a signal-to-noise ratio of up to several orders of magnitude. Examples of such systems in which proximity changes are usefully detected include conformational changes in biomols. resulting from their interaction with their binding partners or ligands. Such biomols. may be, for example, nucleic acids, proteins, peptides, polysaccharides, or other polymeric, naturally-occurring or synthetic mols. These include, by way of non-limiting example, mol. beacons, which detect particular polynucleotide sequences; antibody-antigen interactions, and conformational changes in proteins upon binding to a ligand or substrate. A hairpin loop ssDNA was covalently linked to gold nanoparticles and to different fluorophores and the construct was used in single mismatch detection.

IT **146368-14-1**, Cy5

RL: RCT (Reactant); RACT (Reactant or reagent)
 (methods employing fluorescence quenching by metal surfaces)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HCAPLUS COPYRIGHT 2002 ACS L68 ANSWER 2 OF 18 2001:767507 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

135:315603

TITLE:

SOURCE:

DNA conjugate with functional substance

INVENTOR(S):

Musha, Kiyoshi; Saito, Masako; Ikeshima, Tetsuya

PATENT ASSIGNEE(S):

Asahi Denka Kogyo K. K., Japan Jpn. Kokai Tokkyo Koho, 12 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2001294597	A2	20011023	JP 2000-110909	20000412

AΒ A method is provided for prepn. of a functional substance-incorporated DNA complex. Substances absorbing UV light, visible light, near IR light, or chromophore are incorporated. Fluorescence or phosphorescence emitting substances, thermochromism causing substances, chem. reaction causing substances, or conductive substances, are also used. DNA conjugates with bisphenol A, Sirius Yellow GC, benzopurpurine 4B, ADK Stab AQ 20, ADK Stab LA 32, ADK Stab 1413, 1-ethyl-4-[3-(1-ethy 4(1H)quinolinylidene)-1-propenyl]quinolinium iodide, and 2-[7-(1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indol-2ylidene)-1,3,5-heptatrienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-3Hindolium hydroxide, inner salt, trisodium salt, were prepd.

ΙT 121186-51-4D, DNA conjugate with

RL: ARU (Analytical role, unclassified); PEP (Physical, engineering or chemical process); ANST (Analytical study); PROC (Process) (DNA conjugate with functional substance)

121186-51-4 HCAPLUS

RN 3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-2H-indoliumCN indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-5-sulfo-1-(4-sulfobutyl)-, inner salt, trisodium salt (9CI) (CA INDEX NAME)

●3 Na

L68 ANSWER 3 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:729821 HCAPLUS

DOCUMENT NUMBER: 135:254117

TITLE: Determination of DNA binding activity of DNA binding

proteins

INVENTOR(S): Kulesz-Martin, Molly F.; Liu, Yuangang

PATENT ASSIGNEE(S): Health Research, Inc., USA SOURCE: Eur. Pat. Appl., 26 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
EP 1138781 A2 20011004 EP 2001-106806 20010319

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, SI, LT, LV, FI, RO

JP 2001321199 A2 20011120 JP 2001-103067 20010402 PRIORITY APPLN. INFO.: US 2000-539945 A 20000331

A method is described for quantifying a DNA binding protein in a biol. sample. The method may be used in the absence of radioisotopes, and at a sensitivity greater than the commonly used electrophoretic mobility shift assay (EMSA) method previously discussed. In particular, the method includes the steps of: (a) treating the sample to obtain a liq. to be tested for the DNA binding protein, (b) incubating the liq. contg. the DNA binding protein with a DNA, contg. a binding sequence to which DNA binding protein can specifically bind so that DNA binding protein specifically binds to the DNA, (c) sepg. the DNA with bound DNA binding protein from other DNA and proteins in the liq., and (e) quantifying the DNA binding protein without interference from other proteins of similar mol. wt. in the absence of the use of a radioisotope, and at a sensitivity greater than EMSA. The amt. of DNA binding protein may be quantified in several ways, e.g. directly by immunoreaction to the DNA binding protein or indirectly by amplification of DNA specifically bound to the binding protein followed by quantification of the amplified DNA. In a further embodiment of the invention, the total protein in the sample, having at least one specific immunoreactive site in common with the DNA binding protein, is quantified and compared with the quantity of binding protein to give an indication of relative proportions of bound and unbound protein in the total protein. The present invention esp. relates to methods and kits applicable for the quantification of DNA binding activity of DNA binding proteins for use in both clin. and lab. settings. Such quantifications may be used both fo research and for reliable prognostic indicators in many disease, e.g., cancer.

IT **144377-05-9 146397-20-8**, cy3

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (DNA binding activity of DNA binding proteins detn.)

RN 144377-05-9 HCAPLUS

CN 3H-Indolium, 5-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-2-[5-[5-[2-[(2,5-dioxo-1-pyrrolidinyl)oxy]-2-oxoethyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, sodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

RN 146397-20-8 HCAPLUS

3H-Indolium, 1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

L68 .ANSWER 4 OF 18 HCAPLUS COPYRIGHT 2002 ACS 2001:713617 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

135:268132

TITLE:

Microarray-based analysis of polynucleotide sequence

variations using groups of primers immobilized on

solid phase support

INVENTOR(S): Yu, Zailin; Peng, Zaoyuan; Hu, Qianjin

PATENT ASSIGNEE(S): SOURCE:

Mergen Ltd., USA

PCT Int. Appl., 29 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                                KIND
                                         DATE
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                                A2
       WO 2001071041
                                         20010927
                                                               WO 2001-US9165
                                                                                        20010320
                               А3
       WO 2001071041
                                         20020718
            W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
                   VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
             RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
                   BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                 В1
                                                               US 2000-707366 20001106
       US 6376191
                                         20020423
       US 2002086322
                                                               US 2001-8560
                                 Α1
                                         20020704
                                                                                        20011205
                                                           US 2000-191356P P 20000322
PRIORITY APPLN. INFO.:
                                                                                  A2 20001106
                                                           US 2000-707366
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Solid phase polymerase-mediated amplification approaches using immobilized AB primers on a microarray are provided for detecting sequence variations in a target polynucleotide as compared to a ref. sequence. The group of primers is selected to span a particular region of the ref. sequence and comprises at least four sets of primers: a first set that is exactly complementary to the ref. sequence; and three addnl. sets of primers, each of which is identical to the first set of primers but for the nucleotides at 3'-end, which is different in each of the three sets. The invention provides kit, DNA polymerase, dNTP for detection of sequence variations. The invention can be used to detect sequence variation between a target polynucleotide and a ref. sequence, including single or multiple base substitution, deletion or insertions, and other more complex variations. The methods and compns. provided herein are useful for research and clin. applications, particularly for large scale assays of genetic information in biol. samples of interest.

#### IT 146368-16-3

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES

(Cy3, antibody conjugated with; microarray-based anal. of polynucleotide sequence variations using groups of primers immobilized on solid phase support)

RN 146368-16-3 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

L68 ANSWER 5 OF 18 HCAPLUS COPYRIGHT 2002 ACS 2001:636252 HCAPLUS ACCESSION NUMBER:

135:206422 DOCUMENT NUMBER:

Ligase/polymerase method for detecting cytosine TITLE:

methylation in DNA samples INVENTOR(S): Olek, Alexander; Berlin, Kurt Epigenomics A.-G., Germany PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                          KIND DATE
                                                    APPLICATION NO.
                                                                         DATE
                           ____
                                  _____
                                                     _____
                                                  WO 2001-DE749 20010223
     WO 2001062961
                         A1
                                  20010830
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
               YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                           A1 20010906
                                                     DE 2000-10010281 20000225
      DE 10010281
                                                 DE 2000-10010281 A 20000225
PRIORITY APPLN. INFO.:
     The invention relates to a method for detecting 5-methylcytosine in
     genomic DNA samples. Firstly, a genomic DNA is chem. reacted with a
     reagent, such as bisulfite, whereby 5-methylcytosine and cytosine react
     differently. Afterwards, the pretreated DNA is amplified while using a
     polymerase and at least one primer. In the next step, the amplified
     genomic DNA is hybridized to two different oligonucleotide probes. The
     probes hydridizes adjacent to the cytosine with the unknown methylation
     status. One probe is lengthened by at least one nucleotide, the
     lengthening being subject to the methylation status of the resp. cytosine
     in the genomic DNA sample. The lengthened first probe is then ligated to
     the second probe and the ligation product is detected.
TΤ
     146368-16-3D, conjugates with dCTP
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
      (Biological study, unclassified); ANST (Analytical study); BIOL
      (Biological study); PROC (Process); USES (Uses)
          (Cy3; ligase/polymerase method for detecting cytosine methylation in
         DNA samples)
RN
      146368-16-3 HCAPLUS
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3H-Indolium, 2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-CN dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

IT 146368-14-1D, conjugates with dCTP

RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses) (Cy5; ligase/polymerase method for detecting cytosine methylation in DNA samples)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L68 ANSWER 6 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:635827 HCAPLUS

DOCUMENT NUMBER: 135:206416

TITLE: Method for detecting cytosine methylation in DNA

samples

INVENTOR(S):

PATENT ASSIGNEE(S):

SOURCE:

Olek, Alexander; Berlin, Kurt
Epigenomics A.-G., Germany
PCT Int. Appl., 28 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                          KIND DATE
                                                        APPLICATION NO. DATE
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                                                      WO 2001-DE750 20010223
                                     20010830
      WO 2001062064
                            A2
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                 CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
                 HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
                 LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
           SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
      DE 10010280
                              Α1
                                     20010906
                                                        DE 2000-10010280 20000225
                                                        AU 2001-42280
      AU 2001042280
                              Α5
                                     20010903
                                                                               20010223
                                                     DE 2000-10010280 A 20000225
PRIORITY APPLN. INFO.:
                                                                         W 20010223
                                                     WO 2001-DE750
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AB The invention relates to a method for detecting 5-methylcytosine in genomic DNA samples. Firstly, a genomic DNA is chem. reacted with a reagent, such as bisulfite, whereby 5-methylcytosine and cytosine react differently. Afterwards, the pretreated DNA is amplified while using a polymerase and at least one primer. In the next step, the amplified genomic DNA is hybridized to at least one oligonucleotide probe. The probe hydridizes adjacent to the cytosine with the unknown methylation status. The probe is lengthened by at least one nucleotide, whereby the nucleotide carries a detectable tag, and the lengthening is subject to the methylation status of the resp. cytosine in the genomic DNA sample. Thus, labeled ddCTP and (differently) labeled ddTTP may be added to the probe using DNA polymerase. In the following step, the lengthened oligonucleotides are examd. for the presence of the tag.

IT 146368-16-3D, conjugates with dCTP

RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)

(Cy3; method for detecting cytosine methylation in DNA samples)

RN 146368-16-3 HCAPLUS

CN 3H-Indolium, 2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

IT 146368-14-1D, conjugates with dCTP

RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)

(Cy5; method for detecting cytosine methylation in DNA samples)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

L68 ANSWER 7 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:397101 HCAPLUS

DOCUMENT NUMBER:

135:15070

TITLE:

Continuous time-resolved resonance energy-transfer

assay for polynucleic acid polymerases

INVENTOR(S):

Furfine, Eric Steven; Porter, David John Timothy;

Roberts, Grace Brashear

PATENT ASSIGNEE(S): SOURCE:

Glaxo Group Limited, UK PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND					ND	DATE			A	PPLI	CATIO	ON NO	Э.	DATE				
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WO 2001038587			A:	2	2001	0531		WO 2000-US32536 20001129										
WO 2	2001	03858	37	A.	3	2002	0510											
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	
		ΗU,	ID,	IL,	IN,	IS,	JP,	ΚĖ,	KG,	KΡ,	KR,	KΖ,	LC,	LK,	LR,	LS,	LT,	
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NO,	NZ,	PL,	PT,	RO,	RU,	
		SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	
		YU,	ΖA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM					
	RW:	GH,	GM,	ΚE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,	
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	
		ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,	ΤG			
DIEMI	TO TO TO		TATEO						1 01	000	1 (7)	4 O D	-	1000	1100			

PRIORITY APPLN. INFO.: US 1999-167940P P 19991129 A method of detecting polynucleic acid polymerase activity, including DNA

and RNA polymerase activity. The method includes providing a polynucleic acid primer-template complex labeled with a energy-emitting chem. species and a nucleotide labeled with a energy-emitting chem. species; mixing the polynucleic acid primer-template complex and the nucleotide with a sample comprising or suspected to comprise a polynucleic acid polymerase; prior to, contemporaneously with or after the mixing, exposing the labeled polynucleic acid primer-template complex and the labeled nucleotide to radiation of excitation wavelength for one of the energy-emitting chem. species to thereby excite that energy-emitting chem. species; and detecting a signal produced by energy transfer between the excited energy-emitting chem. species and the other energy-emitting chem. species as a result of incorporation of the nucleotide into the polynucleic acid primer-template complex via the activity of the polynucleic acid polymerase, the detection of the signal indicating polynucleic acid polymerase activity in the sample. Candidate compds. can also be identified as modulators of polynucleic acid polymerase activity via the Thus, the use of Eu-labeled primer-template complex and Cy5-labeled UTP for detn. of steady-state IC50 values for nevirapine, delavirdine, and efavirenz with HIV-1 reverse transcriptase was demonstrated.

#### IT 146368-16-3D, conjugates with

nucleotides/oligonucleotides

RL: ARG (Analytical reagent use); PEP (Physical, engineering or chemical process); ANST (Analytical study); PROC (Process); USES (Uses)

(Cy3; continuous time-resolved resonance energy-transfer assay for polynucleic acid polymerases)

RN 146368-16-3 HCAPLUS CN 3H-Indolium, 2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

# IT 146368-14-1D, conjugates with

nucleotides/oligonucleotides

RL: ARG (Analytical reagent use); PEP (Physical, engineering or chemical process); ANST (Analytical study); PROC (Process); USES (Uses) (Cy5; continuous time-resolved resonance energy-transfer assay for polynucleic acid polymerases)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

## IT 102185-03-5D, Cy2, conjugates with

nucleotides/oligonucleotides

RL: ARG (Analytical reagent use); PEP (Physical, engineering or chemical process); ANST (Analytical study); PROC (Process); USES (Uses) (continuous time-resolved resonance energy-transfer assay for polynucleic acid polymerases) RN

102185-03-5 HCAPLUS 3H-Indolium, 2-[2-[2-chloro-3-[(1,3-dihydro-1,3,3-trimethyl-2H-indol-2-CN ylidene)ethylidene]-1-cyclohexen-1-yl]ethenyl]-1,3,3-trimethyl-, perchlorate (9CI) (CA INDEX NAME)

CM 1

CRN 69415-17-4 CMF C32 H36 C1 N2

CM

CRN 14797-73-0 CMF Cl O4

## MAUPIN 09/829,467

=> d ibib abs hitstr 168 8

L68 ANSWER 8 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:182049 HCAPLUS

DOCUMENT NUMBER: 135:299174

TITLE: Fluorescent high-density labeling of DNA: error-free

substitution for a normal nucleotide

AUTHOR(S): Foldes-Papp, Z.; Angerer, B.; Ankenbauer, W.; Rigler,

R.

CORPORATE SOURCE: MBB, Department of Medical Biophysics, Karolinska

Institute, Stockholm, S-17177, Swed.

SOURCE: Journal of Biotechnology (2001), 86(3), 237-253

CODEN: JBITD4; ISSN: 0168-1656

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

The enzymic incorporation of deoxyribonucleoside triphosphates by a thermostable, 3' 5' exonuclease deficient mutant of the Tgo DNA polymerase was studied for PCR-based high-d. labeling of 217-bp 'natural' DNA in which fluorescent-dUTP was substituted completely for the normal dTTP. The amplified DNA carried two different sorts of tethered dye mols. The rhodamine-green was used for internal tagging of the DNA. Since high-d. incorporation of rhodamine-green-X-dUTP led to a substantial redn. (quenching) of the rhodamine-green fluorescence, a second 'high' quantum yield label, Cy5, was inserted via a 5'-tagged primer in order to identify the two-color product. A theor. concept of fluorescence auto- and cross-correlation spectroscopy developed here was applied to quantify the DNA sequence formed in terms of both the no. of two-color fluorescent mols. and the no. of covalently incorporated rhodamine-green-X-dUMP residues. The novel approach allowed to sep. optically the specific DNA product. After complete, exonucleolytic degrdn. of the two-color DNA we detd. 82-88 fluorescent U\* labels incorporated covalently out of 92 max. possible U\* incorporations. The heavily green-labeled DNA was then isolated by preparative mobility-shift electrophoresis, re-amplified in a subsequent PCR with normal deoxyribonucleoside triphosphates, and re-sequenced. By means of this novel methodol. for analyzing base-specific incorporations that was first developed here, we found that all fluorescent nucleotides and the normal nucleotides were incorporated at the correct positions. The detd. labeling efficiency of 0.89-0.96 indicated that a fraction of the substrate analog was not bearing the fluorophore. The results were used to guide developments in single-mol. DNA sequencing. The labeling strategy (principal approach) for PCR-based high-d. tagging of DNA, which included an appropriate thermostable DNA polymerase and a suitable fluorescent dye-dNTP, was developed here.

# IT 146368-14-1

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (Cy5; fluorescent high-d. labeling of DNA)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HO3S 
$$\sim$$
 CH-CH=CH-CH=CH $\sim$  N=  $\sim$  N=

REFERENCE COUNT:

41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

# MAUPIN 09/829,467

=> d ibib abs hitstr 168 9

L68 ANSWER 9 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:85462 HCAPLUS

DOCUMENT NUMBER:

134:142724

TITLE:

Real-time monitoring of de novo nucleic acid

synthesis using fluorescence resonance

energy transfer

INVENTOR(S):

Okamura, Koji; Sase, Ichiro; Kan, Takayuki; Furusawa, Iwao; Sanze, Kazuyuki; Watanabe, Yuichiro; Kawakami,

Shiqeki

PATENT ASSIGNEE(S):

Bunshi Bio Photonics Kenkyusho K. K., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 9 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE --------------\_\_\_\_\_\_ \_\_\_\_\_ JP 2001029072 A2 20010206 JP 1999-203576 19990716

- AB A novel method for real-time monitoring of nucleic acid synthesis in vitro transcription reactions using fluorescence resonance energy transfer (FRET) is presented. Nucleotide monomers labeled with FRET donor fluorophore and acceptor fluorophore are used in in vivo or in vitro transcription reaction. FRET derived fluorescence is detected for detection of de novo nucleic acid synthesis. RNA synthesis in an in vitro transcription reaction was detected using T7 RNA polymerase, Fluorescein-12-UTP or Bodipy FL-14-UTP as donor, and either Bodipy TMR-14-UTP, Bodipy TR-14-UTP, or Texas Red-5-UTP as acceptor. DNA synthesis was detected using reverse transcriptase, Fluorescein-11-dUTP as donor, and Cy5-dUTP as acceptor.
- 146368-14-1D, Cy5, conjugates with dUTP TΤ RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses) (Cy5; real-time monitoring of de novo nucleic acid synthesis

using fluorescence resonance energy transfer)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-(2,5-dioxo-1-pyrrolidiny1)oxy]-6-oxohexy1]-1,3dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HO3S Me Me CH-CH=CH-CH=CH
$$\stackrel{\text{Et}}{\longrightarrow}$$
 N-R Me Me

## MAUPIN 09/829,467

=> d ibib abs hitstr 168 10

L68 ANSWER 10 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:825711 HCAPLUS

DOCUMENT NUMBER: 134:159683

TITLE: Identification of single fluorescently

labelled mononucleotide molecules in

solution by spectrally resolved time-correlated

single-photon counting

AUTHOR(S): Herten, D. P.; Tinnefeld, P.; Sauer, M.

CORPORATE SOURCE: Physikalisch-Chemisches Institut, Universitat

Heidelberg, Heidelberg, 69120, Germany

SOURCE: Applied Physics B: Lasers and Optics (2000), 71(5),

765-771

CODEN: APBOEM; ISSN: 0946-2171

PUBLISHER: Springer-Verlag

DOCUMENT TYPE: Journal LANGUAGE: English

AB We describe a method to identify single dye-labeled mononucleotide mols. in soln. with high classification probability based on confocal microscopy in combination with spectrally and time-resolved fluorescence detection with two detectors. For efficient excitation of the labeled mononucleotide mols. JA133-dUTP, JA169-dUTP, Cy5-dCTP, and JA242-dUTP a short-pulse diode laser emitting at 634 nm with a repetition rate of 64 MHz was applied. The time-resolved fluorescence signals of individual mols. were analyzed and identified by a max. likelihood estimator (MLE). Scatter plots of spectrally and time-resolved fluorescence data demonstrated the existence of four distinct populations with sym. shape. The distributions of each of the mononucleotide conjugates were detd. by fitting a superposition of two independent Gaussians. Taking only those single-mol. bursts which contain more than 50 photon counts, three labeled mononucleotide mols. were identified in soln. by spectrally and time-resolved fluorescence spectroscopy with a probability of correct classification of .apprxeq. 99%.

IT **146368-14-1**, Cy5

RL: ARG (Analytical reagent use); PRP (Properties); ANST (Analytical study); USES (Uses)

(identification of single fluorescently labeled mononucleotide mols. in soln. by spectrally resolved time-correlated single-photon counting)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L68 ANSWER 11 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:707239 HCAPLUS

DOCUMENT NUMBER: 133:283009

TITLE: Fluorescent squaraine dyes for biological applications

and their preparation

INVENTOR(S): West, Richard Martin; Cummins, William Jonathan;

Nairne, Robert James Domett; Bull, Matthew Graham

PATENT ASSIGNEE(S): Amersham Pharmacia Biotech UK Limited, UK

SOURCE: PCT Int. Appl., 81 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.			KI	ND	DATE		APPLICATION NO.						DATE					
	WO	2000	0584	05	A2 20001005				w W	0 20	00-G	- <b>-</b> 3	20000330						
	WO	2000	0584	05	A	3	2001	0201											
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	
															GH,				
			ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	
			LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	
			SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	
			ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM							
		RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	ΤZ,	UG,	ZW,	AT,	BE,	CH,	CY,	DE,	
			DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	
			CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	TG					
	ΑU	2000	0356	80	A.	5	20001016			AU 2000-35680					20000330				
	EΡ	1165	693		A.	2	2002	0102		E	P 20	00-9	1428	4	2000	0330			
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,	
			ΙE,	SI,	LT,	LV,	FI,	RO											
PRIO	RITY	APP	LN.	INFO	. :				]	EP 1	999-	3025	10	A	1999	0331			
									Ţ	WO 2	000-	GB12:	23	W	2000	0330			

OTHER SOURCE(S):

MARPAT 133:283009

GΊ

Ρ

$$W \longrightarrow NR^{1}R^{2}$$
 $M \longrightarrow NR^{1}R^{2}$ 
 $M \longrightarrow NR^{1}$ 

AB The dyes, with improved photostability, have the structure I or II [R = neg. charge, LG; G = reactive group for bonding to biomols., spectrum- or soly.-modifying group; L = linking group; R1 = H, C1-20 hydrocarbyl, LG; R2 = H, C1-20 hydrocarbyl, OR5, NR6R7, COR7, NR6COR7, N:R8, CO2R7, LG, or NR1R2 = heterocyclyl; R5-R8 = H, C1-20 hydrocarbyl, LG; W = ring system contg. quaternary N conjugated with the cyclobutene ring (via a linking group); m = 1-3] or are (homo- or hetero-) dimers or oligomers thereof. Thus, di-Me squarate was condensed with 1,3,3-trimethyl-2-

methyleneindoline and the resulting methoxycyclobutene compd. was hydrolyzed to the hydroxy analog, which was treated with piperidine to give a I (R = neg. charge, NR1R2 = piperidino, m = 1), absorption .lambda.max 470 nm, fluorescence .lambda.max 495 nm.

IT 299207-42-4

RL: RCT (Reactant); RACT (Reactant or reagent) (prepn. of fluorescent squaraine dyes for biol. applications)

RN 299207-42-4 HCAPLUS

CN 3H-Indolium, 2-[3-[1-(3-aminopropyl)-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

IT 299425-00-6P

RL: SPN (Synthetic preparation); TEM (Technical or engineered material use); PREP (Preparation); USES (Uses)

(prepn. of fluorescent squaraine dyes for biol. applications)

RN 299425-00-6 HCAPLUS

CN Cyclobutenediylium, 1-[(3-ethyl-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene)methyl]-3-[4-[[[3-[2-[3-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1-propenyl]-3,3-dimethyl-3H-indolio]propyl]amino]carbonyl]-1-piperidinyl]-2,4-dihydroxy-, tris(inner salt) (9CI) (CA INDEX NAME)

L68 ANSWER 12 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:441969 HCAPLUS

DOCUMENT NUMBER: 133:86486

TITLE: High throughput assay system using Multi Array Plate

Screen, nuclease protection, oligonucleotide anchors,

APPLICATION NO. DATE

bifunctional linkers, and mass spectrometry

INVENTOR(S):
Kris, Richard M.; Felder, Stephen

PATENT ASSIGNEE(S): USA

SOURCE: PCT Int. Appl., 111 pp.

KIND DATE

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT NO.

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    WO 2000037684 A1 20000629 WO 1999-US30515 19991222
        W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
            CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
            IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
            MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
            SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                       US 1998-218166
PRIORITY APPLN. INFO.:
                                                      Al 19981222
AB
    The present invention relates to compns., app. and methods useful for
    concurrently performing multiple, high throughput, biol. or chem. assays,
    using repeated arrays of probes, called Multi Array Plate Screen (MAPS).
    A combination of the invention comprises a surface, which comprises a
    plurality of test regions, at least two of which, and in a preferred
    embodiment, at least twenty of which, are substantially identical, wherein
    each of the test regions comprises an array of generic anchor mols. The
    anchors are assocd. with bifunctional linker mols., each contg. a portion
    which is specific for at least one of the anchors and a portion which is a
    probe specific for a target of interest. The resulting array of probes is
    used to analyze the presence or test the activity of one or more target
    mols. which specifically interact with the probes. In one embodiment of
    the invention, the test regions (which can be wells) are further
    subdivided into smaller subregions (indentations, or dimples). In one
    embodiment of the invention, ESTs are mapped. In another embodiment, the
    presence of a target nucleic acid is detected by protecting the target
    against nuclease digestion with a polynucleotide fragment, and analyzing
    the protected polynucleotide by mass spectrometry.
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## IT **146368-14-1**, Cy5

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(Cy5, fluorescent probe; high throughput assay system using Multi Array Plate Screen, nuclease protection, oligonucleotide anchors, bifunctional linkers, and mass spectrometry)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

REFERENCE COUNT:

11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L68 ANSWER 13 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:260581 HCAPLUS

DOCUMENT NUMBER: 132:289573

TITLE: Fluorescent probes for chromosomal painting

INVENTOR(S): Cherif, Dorra PATENT ASSIGNEE(S): Genset, Fr.

SOURCE: PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                                 KIND DATE
                                                                 APPLICATION NO. DATE
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                                                                 -----
                                                        WO 1999-FR2517 19991015
                               A1 20000420
       WO 2000022164
             W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ,
                   BY, KG, KZ, MD, RU, TJ, TM
             RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
       FR 2784683
                                 A1 20000421
                                                               FR 1998-12957
                                                                                           19981015
                                                                 AU 1999-60981
       AU 9960981
                                          20000501
                                  Α1
                                                                                           19991015
                                                               EP 1999-947589 19991015
       EP 1121461
                                 A1
                                        20010808
             R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
                   IE, SI, LT, LV, FI, RO
PRIORITY APPLN. INFO.:
                                                             FR 1998-12957
                                                                                      A 19981015
```

WO 1999-FR2517 W 19991015

- The invention concerns fluorescent probes used in multicolor in situ AB fluorescent hybridization methods, and principally chromosomal painting. The probes designed for marking a chromosome are such that they consist of a set of DNA segments more represented in certain chromosomal bands and are obtained by Interspersed Repeated Sequence-PCR amplification from said chromosomes using PCR primers specific for the repeated and dispersed DNA sequences Alu and LINE. The invention further concerns methods for producing said probes, multicolor FISH methods capable of using said probes, and diagnostic kits comprising them. The invention also concerns combinations of fluorophores and optical filters.
- IΤ 146368-16-3DP, Cy3, conjugates with probes RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)
  - (Cy3; fluorescent probes for chromosomal painting)
- RN 146368-16-3 HCAPLUS
- CN 3H-Indolium, 2-[3-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-1-ethyl-3,3dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

ΙT

146368-14-1DP, Cy5, conjugates with probes RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)

(Cy5; fluorescent probes for chromosomal painting)

RN

146368-14-1 HCAPLUS

3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-CN dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HO3S Me Me CH-CH-CH-CH-CH-
$$\frac{Et}{N}$$
 Me Me SO3-

REFERENCE COUNT:

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L68 ANSWER 14 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:219085 HCAPLUS

DOCUMENT NUMBER: 132:247133

TITLE: Optimally fluorescent

oligonucleotides

INVENTOR(S): Nilsen, Thor W.

PATENT ASSIGNEE(S): Poly Probe, Inc., USA

U.S., 14 pp., Cont.-in-part of U.S. Ser. No. 657,961. SOURCE:

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO. DATE
		<b></b>	
US 6046038	A	20000404	US 1997-909539 19970812
US 6072043	A	20000606	US 1996-657961 19960604
US 2002012972	A1	20020131	US 2001-911039 20010723
PRIORITY APPLN. INFO.	:		US 1996-657961 A2 19960604
			US 1997-909539 A1 19970812
			US 2000-482803 B1 20000113

A method is disclosed for the prepn. of optimally fluorescent AΒ oligonucleotides wherein fluorescent dye-

conjugated nucleotide triphosphates are

incorporated into a nucleic acid sequence in a defined repetitive manner which allows for the optimal specific fluorescence of the oligonucleotide. The oligonucleotides of the present invention are useful in the assay of a wide variety of nucleic acid sequences, specifically wherever highly fluorescent nucleic acid probes are desired. The prepn. and purifn. of optimally fluorescent oligonucleotides comprise the steps of (a) prepg. a DNA primer; (b) prepg. a template oligonucleotide contg. a

nucleotide sequence complementary to the primer, and a nucleotide repeat region downstream from the complementary region; (c) annealing the template and primer in a suitable reaction medium contg. a DNA polymerase,

nucleotide triphosphates and fluorescent dye-

conjugated nucleotide triphosphates; (d)

synthesis of complementary strand of the template; (e) attaching the oligonucleotide contg. a target sequence adjacent to the complementary strand; and (f) purifying the optimally fluorescent oligonucleotide. For fluorescence optimization of the

incorporation of dCTP-Cy3, the optimal enzyme is SEQUENASE.RTM. and the optimal spacing is every third mol.

.IT 146397-20-8D, Cy3, conjugate with dCTP

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (prepn. and applications of optimally fluorescent oligonucleotides)

146397-20-8 HCAPLUS RN

CN 3H-Indolium, 1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-2-[3-[1-[6-indolium]][(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## MAUPIN 09/829,467

=> d ibib abs hitstr 168 15

L68 ANSWER 15 OF 18 HCAPLUS COPYRIGHT 2002 ACS

2000:203017 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 132:233995

TITLE: Method of sequencing nucleic acids by applying a

computer alignment algorithm to electrophoretic separation patterns of dideoxy-terminated fragment

mixtures

INVENTOR(S): Mccormick, Randy M.; Briggs, Jonathan

PATENT ASSIGNEE(S): Aclara Biosciences, USA

SOURCE: U.S., 29 pp., Cont.-in-part of U.S. 636,414,

> abandoned. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

DATE PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_\_\_ \_\_\_\_ -----\_\_\_\_\_ US 6043036 A US 1997-977931 19971124 US 1996-636414 19960423 20000328 PRIORITY APPLN. INFO.:

- The present invention describes a method of sequencing nucleic acids in which mixts. of oligonucleotide fragments are derived from sequencing reactions using combinations of the 2',3'-dideoxynucleoside 5'triphosphate or 3' deoxynucleoside 5'-triphosphate terminators and appropriate concns. of four dNTPs (2'-deoxynucleoside 5' triphosphates, e.g., dATP, dCTP, dGTP, dTTP, dITP, 7-deaza-GTP). These fragments are generated by enzymic extension of a primer hybridized to the single-stranded template DNA to be sequenced. In contrast to common slab gel sequencing methods, the method of the instant invention does not require precise alignment of the four sepn. sets of the terminated fragments to permit deduction of the DNA sequence. Instead the relative positions of the nucleotide bases in sep. mixts. can be deduced from binary-coded sequence string sets corresponding to the presence or absence of particular fragments by applying a computer alignment algorithm. In addn., the method possesses inherent redundancy in the sepns., which facilitates sequence assignment by resolving sequence uncertainties or anomalies. The method was applied to the detn. of the known sequence of M13mp18 DNA, and a lengthy stretch of sequence in the middle was correctly detd.
- 146368-14-1D, FluoroLink Mono Reactive Dye Cy5, conjugate ΙT with sequencing primers 146368-16-3D, FluoroLink Mono Reactive Dye Cy3, conjugate with sequencing primers
  - RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (method of sequencing nucleic acids by applying a computer alignment algorithm to electrophoretic sepn. patterns of dideoxy-terminated fragment mixts.)
- 146368-14-1 HCAPLUS RN
- 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-CN dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HO3S 
$$N-R$$
  $CH-CH-CH-CH-CH-CH-N+$   $N-R$   $Me$   $Me$   $SO_3-$ 

RN

 $\begin{array}{lll} 146368-16-3 & \text{HCAPLUS} \\ 3\text{H-Indolium, } 2-[3-[1-[6-[(2,5-\text{dioxo-1-pyrrolidinyl})\,\text{oxy}]-6-\text{oxohexyl}]-1,3-\text{dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene}]-1-\text{propenyl}]-1-\text{ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI)} \\ & (\text{CA INDEX NAME}) \\ \end{array}$ CN

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L68 ANSWER 16 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1996:637657 HCAPLUS

DOCUMENT NUMBER:

125:321156

TITLE:

Hybridization of polynucleotides conjugated with chromophores and fluorophores to generate

donor-to donor energy transfer system

INVENTOR(S):
PATENT ASSIGNEE(S):

Heller, Michael J. Nanogen, Inc., USA

SOURCE:

U.S., 24 pp., Cont.-in-part of U.S. Ser. No. 790, 262,

abandoned.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 37

PATENT INFORMATION:

	PATENT NO.				KIN		DATE			A	PPI	PLICATION NO.			DATE				
	US	5565	322		А		19961 19930	.015		U	IS 1	1994	1-23	2233	3	1994	0505		
	WO						19930	1212		V	0.1	1992	05	902	′	1992	1100		
			AU,				, DK,	гc	r D	CD	CD	) т	· c·	τm	тп	MC	NIT	C F	
	ГD						20010											3E	
							20010			1	11 2	.000	, 12	. 1 2 / -	,	1))2	1100		
	шт						, DK,		FR	GB	GR	<b>₹</b> Т	т.	T.T.	T.11	NT.	SE	MC	TE
	пс	5849	-	DD,		ונוט	19981											110,	
	US	5835	404		A		20000 19981	110		Ü	is 1	997	'-90	6569	,	1997	0805		
		6067	246		A		20000	523		Ü	IS 1	998	-12	9740	)	1998	0805		
	US	6385	080		В:	l	20020	507		U	is 2	2000	-56	8527	7	2000	0509		
	US	6416	953		В.	1	20020	709		U	IS 2	2000	72	4753	3	2000	1128		
PRIOR																1991			
									1	WO 1	992	2-US	982	:7	W	1992	1106		
										EP 1	992	2-92	2522	:5	A3	1992	1106	*	
									1	US 1	993	3-14	650	4	A2	1993	1101		
																1994			
																1994			
												-				1994			
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																1994			
													-			1995			
										US 1						1996			
																1997			
																1998			
7) D	mı.						conte	,						_		1998			.1

AB The presence invention contemplates chromophore-contg. polynucleotides having at least two donor chromophores operatively linked to the polynucleotide by linker arms, such that the chromophores are positioned by linkage along the length of the polynucleotide at a donor-donor transfer distance, and at least one fluorescing acceptor chromophore operatively linked to the polynucleotide by a linker arm, such that the fluorescing acceptor chromophore is positioned by linkage at a donor-acceptor transfer distance from at least one of the donor chromophores, to form a photonic structure for collecting photonic energy and transferring the energy to an acceptor chromophore. A 14 nm photonic antenna structure was assembled from 4 oligonucleotides: a 16-mer acceptor unit 5.4 nm long labeled with sulforhodamine 101, a 30-mer intermediate

donor unit 10.2 nm long labeled with 2 **fluoresceins** sepd. by 6 **nucleotides** (or 2.4 nm), a 29-mer intermediate donor unit 9.9 nm long labeled with 2 **fluoresceins** sepd. by 6 **nucleotides**, and a terminal donor 15-mer 5.1 nm long labeled with a single fluorescein. When illuminated with 495 nm light, energy transfer to the acceptor unit was about 76% efficient. 30% Of the transferred energy was found to come from the intermediate oligonucleotide which has its first donor group located 6.8 nm from the acceptor group, i.e. a distance well beyond the Foerster distance. This energy transfer is abolished when the structure is destroyed by heating to 90.degree.

IT 54849-69-3, IR144

RL: NUU (Other use, unclassified); USES (Uses)
 (acceptor chromophore, conjugate with oligonucleotide;
 hybridization of polynucleotides conjugated with chromophores
 and fluorophores to generate donor-to donor energy transfer system)

54849-69-3 HCAPLUS 1H-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-1,1-dimethyl-3-(3-sulfopropyl)-

2H-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-1,1-dimethyl-3-(3-sulfopropyl)-2H-benz[e]indol-2-ylidene]ethylidene]-2-[4-(ethoxycarbonyl)-1-piperazinyl]-1-cyclopenten-1-yl]ethenyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

RN

CN

CRN 55660-40-7 CMF C50 H58 N4 O8 S2

CM 2

CRN 121-44-8 CMF C6 H15 N

L68 ANSWER 17 OF 18 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1995:3281 HCAPLUS

DOCUMENT NUMBER: 122:155746

TITLE: Fluorescent dyes for labeling biosubstances for

analysis

INVENTOR(S): Katayose, Mitsuo; Tai, Seiji; Watanabe, Hiroo

PATENT ASSIGNEE(S): Hitachi Chemical Co Ltd, Japan SOURCE: Jpn. Kokai Tokkyo Koho, 8 pp.

CODEN: JKXXAF

DOCUMENT TYPE: LANGUAGE: Patent Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

JP 05287209 A2 19931102 JP 1992-88743 19920409

OTHER SOURCE(S): MARPAT 122:155746

GI For diagram(s), see printed CA Issue.

The fluorescent dyes I (A1, A2 = benzene, naphthalene, etc.; R1-3 = H, alkyl, alkoxy; X1 = S, O, etc.; X2 = S, O, CO, etc.; and L = polymethylene) and biosubstance (e.g. vitamin, nucleotide, or protein) labeled with the fluorescent dye are prepd. and used for antigen, pharmaceutical, or DNA anal. Four such fluorescent dyes, their sulfonylchloride derivs. and p-aminobenzoic acid adducts, and 3-(4-aminobutyl)morphine labeled with these dyes were prepd. for morphine anal. A labeled DNA probe, i.e. GTTTCCCAGTCACGAC, was also prepd. for DNA sequence anal. The detection of the dye is not affected by heme in blood and is therefore a good test reagent for clin. anal.

IT 154187-62-9P 161159-47-3P 161159-48-4P

RN 154187-62-9 HCAPLUS

CN Quinolinium, 4-[5-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-, inner salt (9CI) (CA INDEX NAME)

RN 161159-47-3 HCAPLUS

CN 3H-Indolium, 3,3-dimethyl-2-[2-[3-(ar-methyl-4H-1-benzopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]-1-(3-sulfopropyl)-, inner salt (9CI) (CA INDEX NAME)

D1-Me

RN 161159-48-4 HCAPLUS

CN 3H-Benz[g]indolium, 3,3-dimethyl-2-[2-[3-(methyl-4H-1-benzothiopyran-4-ylidene)-1-cyclopenten-1-yl]ethenyl]sulfo-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

D1-SO3H

D1-Me

RN 154187-62-9 HCAPLUS

CN Quinolinium, 4-[5-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-, inner salt (9CI) (CA INDEX NAME)

## => d ibib abs hitstr 168 18

L68 ANSWER 18 OF 18 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1993:642929 HCAPLUS

DOCUMENT NUMBER: 119:242929

TITLE:

Polynucleotides conjugated with chromophores

and fluorophores for determination of nucleic acid

INVENTOR(S): Heller, Michael J. PATENT ASSIGNEE(S): Nanotronics, Inc., USA SOURCE: PCT Int. Appl., 83 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 37

PATENT INFORMATION:

	TENT NO.		ND DATE	APPLICATION NO. DATE
	9309128		1 19930513	WO 1992-US9827 19921106
	RW: AT, 9331364	BE, CH,	DE, DK, ES, 1 19930607	FR, GB, GR, IE, IT, LU, MC, NL, SE AU 1993-31364 19921106
EP	620822	A	2 19960328 1 19941026 1 20010530	EP 1992-925225 19921106
JP	R: AT, 07502992	BE, CH,	DE, DK, ES, 2 19950330	FR, GB, GR, IE, IT, LI, LU, MC, NL, SE JP 1992-508793 19921106 EP 2000-121275 19921106
EP	1067134 R: AT,	A BE, CH,	3 20010502 DE, DK, ES,	P. FR, GB, GR, IT, LI, LU, NL, SE, MC, IE
US	5565322	A	19961015	ES 1992-925225 19921106 US 1994-232233 19940505 US 2000-724753 20001128
PRIORIT	Y APPLN.	INFO.:		US 1991-790262 A2 19911107 EP 1992-925225 A3 19921106 WO 1992-US9827 A 19921106 US 1994-232233 A1 19940505
				US 1994-250951 A2 19940527 US 1998-123638 A1 19980728

A method for detn. of a nucleic acid of interest with a photonic energy AΒ transfer system using a polynucleotide labeled with .gtoreg.2 (non) fluorescing donor chromophores at a donor-donor transfer distance and a fluorescing acceptor chromophore at a donor-acceptor distance. Alternatively, the fluorescing acceptor chromophore is located on a different polynucleotide. The method comprises mixing of the (non) fluorescing donors and fluorescing acceptor-labeled polynucleotide, which contained a complementary sequence to the nucleic acid of interest, with a nucleic acid sample; hybridizing; exciting the donor (non)fluorescing chromophore; and detecting the presence of photonic energy transfer.

#### ΙT 54849-69-3

RL: ANST (Analytical study)

(as donor chromophore, in labeling polynucleotides for detn. of nucleic acid by photonic energy transfer)

54849-69-3 HCAPLUS RN

1H-Benz[e]indolium, 2-[2-[3-[[1,3-dihydro-1,1-dimethyl-3-(3-sulfopropyl)-CN 2H-benz[e]indol-2-ylidene]ethylidene]-2-[4-(ethoxycarbonyl)-1-piperazinyl]-1-cyclopenten-1-yl]ethenyl]-1,1-dimethyl-3-(3-sulfopropyl)-, inner salt, compd. with N, N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CRN 55660-40-7 CMF C50 H58 N4 O8 S2

CM 2

CRN 121-44-8

CMF C6 H15 N

Jennich sensch

## MAUPIN 09/829,467

```
=> d gue 169
L1
             78 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  SHINOKI H?/AU
L2
            360 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  INOMATA H?/AU
L3
           2835 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  KOJIMA M?/AU
L4
            514 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  SUDO Y?/AU
L5
            112 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  SESHIMOTO O?/AU
L6
           3873 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  (L1 OR L2 OR L3 OR L4 OR L5)
1.7
             70 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  L6 AND FLUORESCEN?
L8
              6 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  L7 AND NUCLEOTID?
L9
          89880 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  ?CYANIN? OR ?STYRYL?
L10
              1 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                 L8 AND L9
L11
             53 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
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                BI OR 122-51-0/BI OR 130536-69-5/BI OR 14315-97-0/BI OR
                146-91-8/BI OR 1927-31-7/BI OR 20309-92-6/BI OR 2056-98-6/BI
                OR 23065-05-6/BI OR 2564-35-4/BI OR 25981-83-3/BI OR 327174-86-
                7/BI OR 3590-36-1/BI OR 365-07-1/BI OR 365-08-2/BI OR 366451-16
                -3/BI OR 366451-17-4/BI OR 366451-18-5/BI OR 366451-19-6/BI OR
                366451-20-9/BI OR 366451-21-0/BI OR 366451-22-1/BI OR 366451-23
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                366451-27-6/BI OR 366451-28-7/BI OR 39923-67-6/BI OR 39923-68-7
                /BI OR 4224-70-8/BI OR 42467-24-3/BI OR 491-97-4/BI OR
                52940-67-7/BI OR 52988-98-4/BI OR 56-65-5/BI OR 58-64-0/BI OR
                58-97-9/BI OR 58-98-0/BI OR 61-19-8/BI OR 62306-05-2/BI OR
                628-89-7/BI OR 63-37-6/BI OR 63-38-7/BI OR 63-39-8/BI OR
                65-47-4/BI OR 75-03-6/BI OR 76528-21-7/BI OR 80677-38-9/BI OR
                85-32-5/BI OR 86-01-1/BI OR 95-50-1/BI)
L12
              1 SEA FILE=HCAPLUS ABB=ON
                                                 L10 AND L11
                                         PLU=ON
L15
          16673 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  ?STYRYL?
L16
         863910 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  (DNA OR ?NUCLEOTID? OR
                NUCLEIC)
L18
          18654 SEA FILE=HCAPLUS ABB=ON
                                         PLU=ON
                                                  (SULFONATES/CT OR SULFONATION/
                CT OR "SULFONIC ACIDS"/CT)
L19
           6154 SEA FILE=HCAPLUS ABB=ON PLU=ON "PHOSPHATES, BIOLOGICAL
                STUDIES"/CT
L34
                STR
                13
                .G2√ Cy
                             Me√C√√Me
                             10 @11 12
VAR G1=0/S/11
REP G2=(2-9) C
NODE ATTRIBUTES:
CONNECT IS E3 RC AT
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED
GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 14
STEREO ATTRIBUTES: NONE
L36
          45490 SEA FILE=REGISTRY SSS FUL L34
L39
           4949 SEA FILE=REGISTRY ABB=ON PLU=ON
                                                  L36 AND ?SULFON?/CNS
L40
            639 SEA FILE=REGISTRY ABB=ON PLU=ON
                                                  L36 AND ?PHOSPH?/CNS
L41
          39918 SEA FILE=REGISTRY ABB=ON PLU=ON L36 NOT (L39 OR L40)
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# MAUPIN 09/829,467

L42	561	SEA	FILE=REGISTRY ABB=ON	I PLU=ON	L41 AND OC4/ES
L46	8330	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L16(3A) FLUORES?
L50	68099	SEA	FILE=HCAPLUS ABB=ON	PLU≃ON	L15 OR ?CYANINE?
L61	39357	SEA	FILE=REGISTRY ABB=ON	N PLU≔ON	L41 NOT L42
L62	15024	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L61
L63	266	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L46 AND L62
L64	74	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L63 AND ?CONJUGAT?
L65	21	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L64 AND L50
L66	26	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L64 AND ((L18 OR L19) OR
		?SUI	LFON? OR ?PHOSPH?)		
L67	8	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L65 AND L66
L69	7	SEA	FILE=HCAPLUS ABB=ON	PLU=ON	L67 NOT L12

## => d ibib abs hitstr 169 1

L69 ANSWER 1 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:256388 HCAPLUS

DOCUMENT NUMBER: 136:296159

TITLE: Modified carbocyanine dyes and their

conjugates and their uses

INVENTOR(S): Leung, Wai-Yee; Cheung, Ching-Ying; Yue, Stephen

PATENT ASSIGNEE(S): Molecular Probes, Inc., USA

SOURCE: PCT Int. Appl., 107 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
APPLICATION NO. DATE
        PATENT NO.
                              KIND DATE
                                                                  WO 2001-US30404 20010928
        WO 2002026891
                                   A1
                                               20020404
               W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                      CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                               20020620
                                                                         US 2001-968401
        US 2002077487
                                      A1
                                                                                                      20010929
                                                                         US 2001-969853
        US 2002064794
                                      Α1
                                               20020530
                                                                                                       20011001
                                                                    US 2000-236637P P 20000929
PRIORITY APPLN. INFO.:
                                                                    US 2001-276870P P 20010316
```

OTHER SOURCE(S): MARPAT 136:296159

AB Chem. reactive carbocyanine dyes incorporating an indolium ring moiety that is substituted at the 3-position by a reactive group or by a conjugated substance are disclosed. Conjugation through this position results in spectral properties that are uniformly superior to those of conjugates of spectrally similar dyes wherein attachment is at a different position. The invention includes deriv. compds. having one or more benzo nitrogens. The fluorescent dyes have a reduced tendency to self-assoc. and may be used for staining biol. samples.

IT 407627-63-8P 407627-69-4P 407627-86-5P 407627-90-1P 407628-25-5P

RL: BUU (Biological use, unclassified); IMF (Industrial manufacture); BIOL (Biological study); PREP (Preparation); USES (Uses)

(dye; prodn. of indolium fluorescent cyanine dyes and their biol. use)

RN 407627-63-8 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-(6-hydrazino-6-oxohexyl)-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-62-7

CMF C36 H48 N4 O13 S4

CRN 121-44-8 CMF C6 H15 N

RN 407627-69-4 HCAPLUS

CN 3H-Indolium, 2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1-propenyl]-3-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-68-3 CMF C38 H47 N3 O16 S4

CM 2

CRN 121-44-8 CMF C6 H15 N

RN

407627-86-5 HCAPLUS 3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-CN indol-2-ylidene]-1,3,5-heptatrienyl]-3-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N, N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM1

CRN 407627-85-4

CMF C42 H51 N3 O16 S4

CM2

CRN 121-44-8 CMF C6 H15 N

RN 407627-90-1 HCAPLUS

CN 3H-Indolium, 2-[5-(1,3-dihydro-1,3,3-trimethyl-5-sulfo-2H-indol-2-ylidene)-1, 3-pentadienyl] -3-[6-[(2, 5-dioxo-1-pyrrolidinyl)oxy] -6-oxohexyl]  $-\overline{1}$ , 3dimethyl-5-sulfo-, inner salt, compd. with N, N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM1

CRN 407627-89-8

CMF C36 H41 N3 O10 S2

CRN 121-44-8 CMF C6 H15 N

RN 407628-25-5 HCAPLUS

CN 3H-Pyrrolo[2,3-b]pyridinium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-7-(3-sulfopropyl)-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

K

# IT 407627-61-6P

RL: BUU (Biological use, unclassified); IMF (Industrial manufacture); RCT (Reactant); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(dye; prodn. of indolium fluorescent cyanine dyes and their biol. use)

RN 407627-61-6 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-60-5 CMF C40 H49 N3 O16 S4

$$CH=CH=CH=CH-CH=N$$
 $N^{+}$ 
 $CH=CH-CH=CH-CH=N$ 
 $Me$ 
 $Me$ 
 $SO_{3}H$ 

CRN 121-44-8 CMF C6 H15 N

#### 407627-59-2P 407627-67-2P 407627-73-0P IT 407627-84-3P 407627-88-7P

RL: IMF (Industrial manufacture); RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)

(intermediate; prodn. of indolium fluorescent cyanine dyes and their biol. use)

RN

407627-59-2 HCAPLUS 3H-Indolium, 3-(5-carboxypentyl)-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-CN (3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-methyl-5-sulfo-1-(3sulfopropyl)-, inner salt, compd. with N, N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM1

407627-58-1 CRN CMF C36 H46 N2 O14 S4

$$O_3S - (CH_2)_3$$
  $(CH_2)_3 - SO_3H$   $(CH_2)_3 - SO_3H$ 

CRN 121-44-8 CMF C6 H15 N

RN 407627-67-2 HCAPLUS

CN 3H-Indolium, 3-(5-carboxypentyl)-2-[3-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1-propenyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-66-1 CMF C34 H44 N2 O14 S4

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 407627-73-0 HCAPLUS

CN 3H-Indolium, 3-(5-carboxypentyl)-2-[5-(1,3-dihydro-1,3,3-trimethyl-5-sulfo-2H-indol-2-ylidene)-1,3-pentadienyl]-1,3-dimethyl-5-sulfo-, inner salt, compd. with N,N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-72-9 CMF C32 H38 N2 O8 S2

CRN 121-44-8 CMF C6 H15 N

RN 407627-84-3 HCAPLUS

CN 3H-Indolium, 3-(5-carboxypentyl)-2-[7-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-83-2 CMF C38 H48 N2 O14 S4

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 407627-88-7 HCAPLUS

CN 3H-Indolium, 3-(5-carboxypentyl)-2-[5-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-methyl-1-(3-

sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-87-6 CMF C36 H46 N2 O8 S2

CM 2

CRN 121-44-8 CMF C6 H15 N

IT 407627-65-0P 407627-71-8P 407627-75-2P 407627-77-4P 407627-79-6P 407627-81-0P 407627-92-3P 407627-94-5P 407628-29-9P

RL: IMF (Industrial manufacture); PREP (Preparation) (prodn. of indolium fluorescent cyanine dyes and their biol. use)

RN 407627-65-0 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-[6-[[2-(2,5-dihydro-2,5-dioxo-1H-pyrrol-1-yl)ethyl]amino]-6-oxohexyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-64-9 CMF C42 H52 N4 O15 S4

CRN 121-44-8 CMF C6 H15 N

RN 407627-71-8 HCAPLUS

CN 3H-Indolium, 3-(5-carboxypentyl)-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,3-dimethyl-5-sulfo-, inner salt, compd. with N,N-diethylethanamine (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-70-7 CMF C34 H42 N2 O11 S3

Me (CH2) 
$$3-SO_3H$$

N+ CH= CH- CH= CH- CH

Me Me SO\_3H

HO\_2C- (CH2) 5 Me

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 407627-75-2 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-(6-hydroxyhexyl)-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-74-1

CMF C36 H48 N2 O13 S4

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 407627-77-4 HCAPLUS

CN 3H-Indolium, 3-(6-aminohexyl)-2-[5-[1,3-dihydro-3,3-dimethyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-methyl-5-sulfo-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-76-3

CMF C36 H49 N3 O12 S4

CRN 121-44-8 CMF C6 H15 N

RN 407627-79-6 HCAPLUS

CN Benzothiazolium, 2-[5-[3-(5-carboxypentyl)-1,3-dihydro-3-methyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:2) (9CI) (CA INDEX NAME)

CM :

CRN 407627-78-5

CMF C33 H40 N2 O11 S4

CM 2

CRN 121-44-8 CMF C6 H15 N

RN 407627-81-0 HCAPLUS

CN Benzothiazolium, 2-[5-[3-(5-carboxypentyl)-1,3-dihydro-3-methyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-6-sulfo-3-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:3) (9CI) (CA INDEX NAME)

CRN 407627-80-9

CMF C33 H40 N2 O14 S5

$$HO_3S - (CH_2)_3$$
  $(CH_2)_3 - SO_3^ N$ 
 $CH - CH = CH - CH = CH$ 
 $N^+$ 
 $HO_3S$ 
 $HO_2C - (CH_2)_5$ 

CM 2

CRN 121-44-8 CMF C6 H15 N

Et | Et- N- Et

RN 407627-92-3 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3-methyl-1-(3-sulfopropyl)-, inner salt, compd. with N,N-diethylethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 407627-91-2

CMF C40 H49 N3 O10 S2

CRN 121-44-8 CMF C6 H15 N

RN 407627-94-5 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[3-(5-carboxypentyl)-1,3-dihydro-3-methyl-5-sulfo-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-6,8-disulfo-3-(3-sulfopropyl)-, inner salt (9CI) (CA INDEX NAME)

RN 407628-29-9 HCAPLUS

CN 3H-Pyrrolo[2,3-b]pyridinium, 2-[6-methoxy-2-[(3-methyl-6-sulfo-2(3H)-benzothiazolylidene)methyl]-6-oxo-1-hexenyl]-3,3,7-trimethyl-, inner salt (9CI) (CA INDEX NAME)

# IT 407628-24-4P

RL: IMF (Industrial manufacture); RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)

(prodn. of indolium fluorescent **cyanine** dyes and their biol. use)

RN .407628-24-4 HCAPLUS

CN 3H-Pyrrolo[2,3-b]pyridinium, 2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-7-(3-sulfopropyl)-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

K

REFERENCE COUNT:

5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Page 17

### => d ibib abs hitstr 169 2

L69 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:136073 HCAPLUS

DOCUMENT NUMBER: 136:196564

TITLE: Non-fluorescent asymmetric cyanine dye

compounds useful for quenching reporter dyes

INVENTOR(S): Lee, Linda G.; Graham, Ronald J.; Mullah, Khairuzzaman

B.; Haxo, Francis T.

PATENT ASSIGNEE(S): PE Corporation (NY), USA

SOURCE: U.S., 62 pp., Cont.-in-part of U.S. 6,080,868.

CODEN: USXXAM
DOCUMENT TYPE: Patent

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATE	NT NO.	KIND	DATE		APPLICATION NO		DATE
						-	
US 6	348596	B1	20020219		US 1999-357740		19990720
US 6	080868	A	20000627		US 1998-12525		19980123
RIORITY	APPLN.	INFO.:	τ	US	1998-12525	Α2	19980123
MILED COLL	D 0 - 1 0 1						

OTHER SOURCE(S): MARPAT 136:196564

GΙ

$$R^{4}$$
 $X$ 
 $CY^{1} = CH)_{p} (CH = CH)_{n}CH = Z$ 
 $R^{5}$ 
 $R^{6}$ 
 $R^{1}$ 

- AB The invention provides asym. cyanine dye compds. I, including substituted forms thereof, which are non-fluorescent quencher mols. The invention further provides reporter-quencher dye pairs, wherein the asym. cyanine dyes are the quenchers, polynucleotides incorporating the asym. cyanine dyes, and nucleic acid hybridization detection methods utilizing the dye-labeled polynucleotides. Nitrothiazole blue XXXIV was prepd. from 2-methylbenzothiazole and used as a quencher dye paired with FAM or TET reporter dyes in Taqman assays.
- IT 235432-77-6D, salts or protected compds. 235432-78-7D, salts or protected compds. 400090-45-1D, salts or protected compds. 400090-51-9
  - RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (as nonfluorescent asym. cyanine dye; non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes)
- RN 235432-77-6 HCAPLUS
- CN Naphtho[2,1-d]thiazolium, 2-[3-[1-(5-carboxypentyl)-4(1H)-quinolinylidene]-1-propenyl]-3-methyl-6,8-dinitro-(9CI) (CA INDEX NAME)

RN

235432-78-7 HCAPLUS
Quinolinium, 1-[3-[(3-carboxypropyl)dimethylammonio]propyl]-4-[3-(3-methyl-CN 6-nitro-2(3H)-benzothiazolylidene)-1-propenyl]- (9CI) (CA INDEX NAME)

RN 400090-45-1 HCAPLUS

CN Benzothiazolium, 2-[3-[1-(5-carboxypentyl)-4(1H)-quinolinylidene]-1propenyl]-3-methyl-6-nitro- (9CI) (CA INDEX NAME)

RN 400090-48-4 HCAPLUS

CN Naphtho[2,1-d]thiazolium, 2-[3-[1-(5-carboxypentyl)-4(1H)-quinolinylidene]-1-propenyl]-3-methyl-6,8-dinitro-, inner salt (9CI) (CA INDEX NAME)

RN 400090-51-9 HCAPLUS

CN Quinolinium, 1-[3-[(3-carboxypropyl)dimethylammonio]propyl]-4-[3-(3-methyl-6-nitro-2(3H)-benzothiazolylidene)-1-propenyl]-, mono(inner salt) (9CI) (CA INDEX NAME)

#### 400090-56-4P ΙT

RL: SPN (Synthetic preparation); PREP (Preparation) (non-fluorescent asym. cyanine dye compds. useful for quenching reporter dyes) 400090-56-4 HCAPLUS

RN

 $\texttt{Benzothiazolium, 2-[3-[1-[6-[(2,5-\texttt{dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1}}$ CN4(1H)-quinolinylidene]-1-propenyl]-3-methyl-6-nitro- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

REFERENCE COUNT:

THERE ARE 84 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## => d ibib abs hitstr 169 3

L69 ANSWER 3 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:310508 HCAPLUS

DOCUMENT NUMBER:

134:323136

TITLE:

Cyanine dyes as labeling reagents for

detection of biological and other materials by

luminescence methods

INVENTOR(S): PATENT ASSIGNEE(S): Waggoner, Alan S.

SOURCE:

Carnegie Mellon University, USA U.S., 20 pp., Cont.-in-part of U.S. 5,627,027.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

US 6225050 B1 20010501 US 1996-745712 19961112 US 5268486 A 19931207 US 1992-884636 19920515 US 5627027 A 19970506 US 1992-831759 19920922 US 5486616 A 19960123 US 1993-158952 19931129 US 5569766 A 19961029 US 1993-158953 19931129 US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO.: US 1986-854347 B1 19860418 US 1992-831759 A2 19920922
US 5268486 A 19931207 US 1992-884636 19920515 US 5627027 A 19970506 US 1992-831759 19920922 US 5486616 A 19960123 US 1993-158952 19931129 US 5569766 A 19961029 US 1993-158953 19931129 US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO:: US 1986-854347 B1 19860418
US 5627027 A 19970506 US 1992-831759 19920922 US 5486616 A 19960123 US 1993-158952 19931129 US 5569766 A 19961029 US 1993-158953 19931129 US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO:: US 1986-854347 B1 19860418
US 5486616 A 19960123 US 1993-158952 19931129 US 5569766 A 19961029 US 1993-158953 19931129 US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO:: US 1986-854347 B1 19860418
US 5569766 A 19961029 US 1993-158953 19931129 US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO:: US 1986-854347 B1 19860418
US 5569587 A 19961029 US 1995-424219 19950419 US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO:: US 1986-854347 B1 19860418
US 6048982 A 20000411 US 1997-873470 19970612 PRIORITY APPLN. INFO.: US 1986-854347 B1 19860418
PRIORITY APPLN. INFO.: US 1986-854347 B1 19860418
IIC 1002_031750
03 1992-031739
US 1988-240756 B1 19880902
US 1992-882802 B1 19920514
US 1992-884636 A3 19920515
US 1996-745712 A3 19961112

MARPAT 134:323136 OTHER SOURCE(S):

Cyanine and related dyes, such as merocyanine,

styryl and oxonol dyes, are strongly light-absorbing and highly luminescent. Cyanine and related dyes having functional groups to make them reactive with amine, hydroxy and sulfhydryl groups are covalently attached to proteins, nucleic acids, carbohydrates, sugars, cells and combinations thereof, and other biol. and nonbiol. materials, to make these materials fluorescent so that they can be detected. The labeled materials can then be used in assays employing excitation light sources and luminescence detectors. For example, fluorescent cyanine and related dyes can be attached to amine, hydroxy or sulfhydryl groups of avidin and to antibodies and to lectins. Thereupon, avidin labeled with cyanine type dyes can be used to quantify biotinylated materials and antibodies conjugated with cyanine-type dyes can be used to detect and measure antigens and haptens. In addn., cyanine-conjugated lectins can be used to detect specific carbohydrate groups. Also, cyanineconjugated fragments of DNA or RNA can be used to identify the presence of complementary nucleotide sequences in DNA or RNA.

IT 336850-22-7P

> RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(as reactive dye, antibody labeling with; cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN336850-22-7 HCAPLUS

CN3H-Indolium, 2-[7-[5-[[(4,6-dichloro-1,3,5-triazin-2-yl)amino]methyl]-1-[3[(2,5-dioxo-l-pyrrolidinyl)oxy]-3-oxopropyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

# IT 336850-18-1P 336850-20-5P 336850-21-6P

RL: SPN (Synthetic preparation); PREP (Preparation)
(as reactive dye; cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN 336850-18-1 HCAPLUS

CN 3H-Indolium, 5-[[(4,6-dichloro-1,3,5-triazin-2-yl)amino]methyl]-2-[5-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-1,3,3-trimethyl-, inner salt (9CI) (CA INDEX NAME)

PAGE 1-B

- so<sub>3</sub>-

RN 336850-20-5 HCAPLUS

CN 3H-Indolium, 2-[3-[1,3-dihydro-1-(3-isothiocyanatopropyl)-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-1,3,3-trimethyl-, bromide (9CI) (CA INDEX NAME)

• Br-

RN 336850-21-6 HCAPLUS

CN 3H-Indolium, 2-[3-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1-propenyl]-5-[(iodoacetyl)amino]-1,3,3-trimethyl-, inner salt (9CI) (CA INDEX NAME)

# IT 120725-04-4P 336850-24-9P 336850-26-1P 336850-28-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(cyanine dyes as labeling reagents for detection of biol. and other materials by luminescence methods)

RN 120725-04-4 HCAPLUS

CN 3H-Indolium, 5-[(chloroacetyl)amino]-2-[3-[1,3-dihydro-3,3-dimethyl-1-(3-sulfopropyl)-2H-indol-2-ylidene]-1-propenyl]-1,3,3-trimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 336850-24-9 HCAPLUS

CN 3H-Indolium, 1-(2-carboxyethyl)-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 336850-26-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1,3-dihydro-1-(5-hydroxypentyl)-3,3-dimethyl-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

Et 
$$(CH_2)_5-OH$$

$$N^+$$

$$CH=CH-CH=CH-CH$$

$$Me$$

$$Me$$

$$Me$$

RN 336850-28-3 HCAPLUS

CN 1H-Benz[e]indolium, 2-[5-[1,3-dihydro-3-(5-hydroxypentyl)-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-3-ethyl-1,1-dimethyl-6,8-disulfo-, inner salt, monopotassium salt (9CI) (CA INDEX NAME)

K

REFERENCE COUNT:

THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## => d ibib abs hitstr 169 4

L69 ANSWER 4 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2000:402040 HCAPLUS

DOCUMENT NUMBER:

133:28254

TITLE:

Detection of biomaterial using polyamide or

polysulfone membrane support and fluorescent-labeled binding agent Dubitsky, Andrew; Decollibus, Damien

INVENTOR(S):

Pall Corporation, USA

PATENT ASSIGNEE(S):

PCT Int. Appl., 39 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

P	PATENT NO.					KIND DATE			A	PPLI	CATI	DATE							
		000034522						WO 1999-US29000 19991206											
W	0 2000	10345	22	A	3	2001	0607												
	W:	ΑE,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	CU,		
						EE,													
		IN, IS,			KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,		
		MG, MK,			MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,		
		SL,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,			
		BY,	KG,	KZ,	MD, RU, TJ,		MT												
	RW	GH,	GM,	ΚE,	LS,	MW,	SD,	SL,	SZ,	ΤZ,	UG,	ΖW,	AT,	BE,	CH,	CY,	DE,		
		DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,		
		CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG						
PRIORI'	IORITY APPLN. INFO.:					1			US 1998-111915P			P 19981211							
									US 1999-392793				Α	A 19990909					
											US 1999-163788P P 19991105								

- AΒ A method and system for detecting a labeled complex including biomaterial without stringency washing after complexing and/or without amplifying the label is disclosed. The method uses a polyamide or polysulfone membrane support and a fluorescent-labeled binding agent. Reverse dot blot assays for a .beta.-globin sequence and protein dot blots for mouse IgG were performed using various membranes and red fluorescent dye-labeled
- ΙT 146368-14-1D, Cy5, end-labeled conjugates with oligonucleotide probes
  - RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (Cy5; detection of biomaterial using polyamide or polysulfone membrane support and fluorescent-labeled binding agent)
- RN 146368-14-1 HCAPLUS
- CN 3H-Indolium, 2-[5-[1-[6-(2,5-dioxo-1-pyrrolidiny1)oxy]-6-oxohexy1]-1,3dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

HO3S

Me Me

$$CH-CH=CH-CH=CH$$
 $N+$ 
 $N N-$ 

IT 251102-88-2D, IRD 700, end-labeled conjugates with

oligonucleotide probes

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (detection of biomaterial using polyamide or **polysulfone** membrane support and fluorescent-labeled binding agent)

RN 251102-88-2 HCAPLUS

CN lH-Benz[e]indolium, 2-[5-[3-(5-carboxypentyl)-1,3-dihydro-1,1-dimethyl-2H-benz[e]indol-2-ylidene]-1,3-pentadienyl]-1,1-dimethyl-3-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

## MAUPIN 09/829,467

## => d ibib abs hitstr 169 5

L69 ANSWER 5 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1999:495475 HCAPLUS

DOCUMENT NUMBER: 131:141745

TITLE: Energy transfer dyes as labels in biological systems

INVENTOR(S): Flick, Parke

PATENT ASSIGNEE(S): Amersham Pharmacia Biotech, Inc., USA

SOURCE: PCT Int. Appl., 31 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	KIND DA		DATE	DATE			PPLI	CATI	ON NO	DATE							
WO	O 9939203 W: AU, CA,			A.TP	1	19990805			W	0 19	99-U	19990202					
		AT,	BE,		CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,
CA	2319	490		A	Ą	1999	0805		C	A 19	99-2	31949	90	1999	0202		
AU	9925	718		Α	1	1999	0816		Αl	J 19	99-2	5718		1999	0202		
EP	1053	472		Α	1	2000	1122		E.	P 19	99-9	05590	)	1999	0202		
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	FI														
JP	2002	5020	34	$\mathbf{T}$	2	2002	0122		J:	P 20	00-5	2960	6	1999	0202		
PRIORIT	Y APP	LN.	INFO	. :					US 19	998-	1811	1	Α	1998	0203		

WO 1999-US2105 W 19990202

OTHER SOURCE(S): MARPAT 131:141745

AB A novel class of energy transfer dyes, their prepn., and their use as labels in biol. systems is disclosed. The dyes are preferably in the form of cassettes which enable their attachment to a variety of biol. materials. The dyes and the reagents that can be made from them offer a wide variety of fluorescent labels with large Stokes' shifts enabling their use in a variety of fluorescence applications over a wide range of the visible spectrum. Prepn. of FAM-Cysteine-linker-ROX energy transfer dye from L-cysteine, 5-iodoacetamidofluorescein, trifluoroacetyl-protected NHS ester of 6-aminocaproic acid and 5'-ROX-NHS ester is described. With excitation at 488 nm, a strong peak was obsd. at 603 nm, characteristic of the ROX emission and indicating excellent energy transfer.

IT 146368-14-1D, Cy5, conjugates 235749-09-4D,

conjugates 235749-10-7D, conjugates

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (energy transfer dye contg.; energy transfer dyes as labels in biol. systems)

RN 146368-14-1 HCAPLUS

CN 3H-Indolium, 2-[5-[1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-ethyl-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

RN 235749-09-4 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[3-(3-ethyl-5-methyl-2(3H)-benzoxazolylidene)-1-propenyl]-5-methyl- (9CI) (CA INDEX NAME)

Me 
$$CH = CH - CH = N$$
  $Me$   $CH_2)$   $5 - CO_2H$ 

RN 235749-10-7 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[3-(1-ethyl-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene)-1-propenyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

REFERENCE COUNT:

8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## => d ibib abs hitstr 169 6

L69 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1998:708965 HCAPLUS

DOCUMENT NUMBER: 129:335785

TITLE: Acid-labile and enzymically cleavable dye

conjugates for diagnosis with near-IR

radiation and for therapy

Licha, Kai; Riefke, Bjoern; Semmler, Wolfhard; INVENTOR(S):

Wrasidlo, Wolfgang

Institut fuer Diagnostikforschung G.m.b.H. an der PATENT ASSIGNEE(S):

Freien Universitaet Berlin, Germany

PCT Int. Appl., 40 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

GI

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	KII	ND	DATE				APP	LIC	ATI	ON NO	ο.	DATE						
WO	9847	538		A	2	1998	1029			WO	199	8-D	E100	1	1998	0402		
WO	9847	538		A.	3	1999	0121											
	W:	ΑU,	CA,	CN,	HU,	JP,	KR,	NO,	US	3								
	RW:	ΑT,	BE,	CH,	CY,	DE,	DK,	ES,	F]	[, F	R,	GB,	GR,	ΙE,	ΙΤ,	LU,	MC,	NL,
		PT,	SE															
DE	1971	7904		A.	1	1998	1029			DE	199	7-1	9717	904	1997	0423		
AU	9879	057		A.	1	1998	1113			AU	199	8-7	9057		1998	0402		
AU	7337	57		B:	2	2001	0524											
EP	9880	60		A.	2	2000	0329			EΡ	199	8-9	2921:	2	1998	0402		
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GE	3, G	R,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	FI															
JP	2001	5215	30	T	2	2001	1106			JP	199	8-5	4471	5	1998	0402		
NO	9905	181		A		1999	1022			NO	199	9-5	181		1999	1022		
PRIORIT	Y APP	LN.	INFO	.:					DE	199	7-1	971	7904	Α	1997	0423		
									WO	199	8-D	E10	01	W	1998	0402		
OTHER SO	OURCE	(S):			MAR	PAT	129:3	3357	85									`

AB Dyes which fluoresce in the near-IR spectral region are provided, the fluorescence of which is quenched by coupling via a cleavable linker to arom. compds. (e.g. dyes, drugs), antibodies, antibody fragments, or other proteins. Cleavage of such a construct in vivo at a target site (e.g. a tumor or focus of inflammation) leads to an increase in near-IR fluorescence, which can be detected even at deep sites owing to the high transparency of tissues to near-IR radiation. Suitable dyes include tetrapyrrole, tetraazapyrrole, xanthine, phenoxazine, phenothiazine, and

esp. polymethine dyes such as cyanine dyes. Drug-dye conjugates in which the therapeutic activity of the drug is masked by coupling to the dye may serve as prodrugs which, after administration, are cleaved at a target site to release the active agent, as well as the fluorescent dye which may act as photosensitizer, at the site. The linker may be acid labile, i.e. cleavable at the low pH characteristic of tumors and sites of bacterial inflammation, or cleavable by enzymes which occur in diseased tissues, e.g. bacterial enzymes. Thus, a cyanine dye, 5-(1-oxoethyl)-1,1'-(4-sulfobutyl)indotricarbocyanine Na salt (I) was prepd. by reaction of 4-hydrazinophenyl Me ketone with 3-methyl-2-butanone followed by 1,4-butanesultone to form 5-(1-oxoethyl)-1-(4-sulfobutyl)-2,3,3-trimethyl-3H-indolenine and further reaction of this compd. with glutaconaldehyde dianil-HCl. Reaction of I with 4-carboxyphenylsulfonylhydrazine followed by N-hydroxysuccinimide and DCCD produced an acid-labile Nhydroxysuccinimidyl ester, which was coupled to anti-melanoma monoclonal antibody 9.2.27; the antibody conjugate had a fluorescence quantum yield of 0.1%.

215114-65-1P 215114-69-5P 215114-70-8P 215114-73-1P 215114-74-2P 215114-75-3P 215114-76-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(acid-labile and enzymically cleavable dye conjugates for diagnosis with near-IR radiation and for therapy)

RN 215114-65-1 HCAPLUS

CN 3H-Indolium, 5-acetyl-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, sodium salt (9CI) (CA INDEX NAME)

● Na

RN 215114-69-5 HCAPLUS

CN 3H-Indolium, 5-[1-[(4-aminobenzoyl)hydrazono]ethyl]-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

Na

PAGE 1-B

RN 215114-70-8 HCAPLUS

CN 3H-Indolium, 5-[1-[[4-(aminomethyl)benzoyl]hydrazono]ethyl]-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

Na

PAGE 1-B

$$= CH - CH$$
Me
Me
Me

RN

215114-73-1 HCAPLUS 3H-Indolium, 5,5'-[(3,9-diethyl-2,4,8,10-tetraoxaspiro[5.5]undecane-3,9-CN diyl)bis(oxy-6,1-hexanediyliminocarbonyl)]bis[2-[7-[1,3-dihydro-3,3dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3dimethyl-1-(4-sulfobutyl)-, bis(inner salt), disodium salt (9CI) (CA INDEX NAME)

●2 Na

PAGE 1-C

RN

215114-74-2 HCAPLUS 3H-Indolium, 5-carboxy-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-CNindol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

Na

215114-75-3 HCAPLUS RN

3H-Indolium, 2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-inCN ylidene] -1, 3,5-heptatrieny $\bar{1}$ ] -5-[1-[4-[[2-7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4sulfobutyl)-3H-indolium-5-yl]carbonyl]amino]methyl]benzoyl]hydrazono]ethyl ]-3,3-dimethyl-1-(4-sulfobutyl)-, bis(inner salt), disodium salt (9CI) (CA INDEX NAME)

PAGE 1-A

**2** Na

PAGE 1-B

PAGE 1-C

RN 215114-76-4 HCAPLUS

CN 3H-Indolium, 5-[[[[4-[[[1-[(2S,4S)-4-[(3-amino-2,3,6-trideoxy-.alpha.-L-lyxo-hexopyranosyl)oxy]-1,2,3,4,6,11-hexahydro-2,5,12-trihydroxy-7-methoxy-2-naphthacenyl]-2-hydroxyethylidene]hydrazino]carbonyl]phenyl]methyl]amino | carbonyl]-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt, monosodium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry unknown.

# PAGE 1-B

## => d ibib abs hitstr 169 7

L69 ANSWER 7 OF 7 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1997:85099 HCAPLUS

DOCUMENT NUMBER:

126:86792

TITLE:

Fluorescent labeling complexes with large stokes

shifts formed by coupling together cyanine

and other fluorochromes capable of resonance energy

transfer

INVENTOR(S):

Waggoner, Alan Stewart; Mujumdar, Swati Ratnakar;

Mujumdar, Ratnakar Balvant

PATENT ASSIGNEE(S):

Carnegie-Mellon University, USA

SOURCE:

Eur. Pat. Appl., 29 pp.

DOCUMENT TYPE:

CODEN: EPXXDW

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA'	TENT NO.		KIND	DATE	APPLICATION NO. DATE
EP	747700		A2	19961211	EP 1996-303879 19960530
EP	747700		A3	19970507	
EP	747700		В1	20011205	
	R: AT	, BE,	CH, DE	, ES, FI,	FR, GB, IT, LI, NL, SE
US	6008373		A	19991228	US 1995-476880 19950607
GB	2301833		A1	19961218	GB 1996-11453 19960530
GB	2301833		В2	19970716	
EP	943918		A1	19990922	EP 1999-110086 19960530
	R: AT	, BE,	CH, DE	, ES, FR,	GB, IT, LI, NL, SE, FI
AT	210292		E	20011215	AT 1996-303879 19960530
CA	2178308		AA	19961208	CA 1996-2178308 19960605
JP	0910482	5	A2	19970422	JP 1996-146333 19960607
JP	2843296		В2	19990106	
US	6130094		A	20001010	US 1998-152009 19980911
PRIORIT	Y APPLN.	INFO	.:		US 1995-476880 A 19950607
					EP 1996-303879 A3 19960530

- AB The present invention provides low-mol.-wt. fluorescent labeling complexes with large wavelength shifts between absorption of one dye in the complex and emission from another dye in the complex. These complexes can be used, for example, for multiparameter fluorescence cell anal. using a single excitation wavelength. The low mol. wt. of the complex permits materials labeled with the complex to penetrate cell structures for use as probes. The labeling complexes are synthesized by covalently attaching through linkers to form donor-acceptor complexes. Resonance energy transfer from an excited donor to fluorescent acceptor provides wavelength shifts up to 300 nm. The fluorescent labeling complexes preferably contain reactive groups for the labeling of functional groups on target compds., such as derivatized oxy and deoxy polynucleic acids, antibodies, enzymes, lipids, carbohydrates, proteins, and other materials. The complexes may contain functional groups permitting covalent reaction with materials contg. reactive groups.
- 185397-56-2DP, reactions products 185397-56-2P IΤ RL: ARG (Analytical reagent use); PRP (Properties); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses) (fluorescent labeling complexes with large Stokes shifts prepn. for cell anal.)
- RN 185397-56-2 HCAPLUS
- CN 3H-Indolium, 5-(aminomethyl)-2-[3-[5-[[[[3-carboxy-4-(6-hydroxy-3-oxo-3H-

xanthen-9-yl)phenyl]amino]thioxomethyl]amino]methyl]-1,3-dihydro-3,3dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

#### PAGE 1-A

HO3S- (CH2) 4 (CH2) 4-SO3-
$$CH-CH=CH$$

$$CH-CH=CH$$

$$Me$$

$$Me$$

$$Me$$

$$CH2$$

$$CH2$$

$$CH2$$

$$CH2$$

$$CH2$$

$$CH3$$

$$CH4$$

PAGE 1-B

— nн<sub>2</sub>

RN 185397-56-2 HCAPLUS

CN 3H-Indolium, 5-(aminomethyl)-2-[3-[5-[[[[[3-carboxy-4-(6-hydroxy-3-oxo-3H-xanthen-9-yl)phenyl]amino]thioxomethyl]amino]methyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

# PAGE 1-B

## — NH<sub>2</sub>

# IT 185397-43-7P 185397-44-8P 185397-46-0P

RL: ARG (Analytical reagent use); RCT (Reactant); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(fluorescent labeling complexes with large Stokes shifts prepn. for cell anal.)

RN 185397-43-7 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-5-[(1,3-dihydro-1,3-dioxo-2H-isoindol-2-yl)methyl]-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 185397-44-8 HCAPLUS

CN 3H-Indolium, 5-(aminomethyl)-1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-

1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-, inner salt (9CI) (CA INDEX NAME)

RN 185397-46-0 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-5-[[[6-[2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]methyl]-1,3-dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo-, bis(inner salt) (9CI) (CA INDEX NAME)

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IT 185397-41-5P 185397-47-1P 185397-48-2P 185397-49-3P 185397-50-6P 185397-51-7P

RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses) (fluorescent labeling complexes with large Stokes shifts prepn. for

cell anal.)

RN 185397-41-5 HCAPLUS

3H-Indolium, 5-[[4-chloro-6-[[2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-3H-indolium-5-yl]amino]-1,3,5-triazin-2-yl]amino]-2-[5-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, bis(inner salt) (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

RN 185397-47-1 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[3-[3-[6-[[[2-[3-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-3H-indolium-5-yl]methyl]amino]-6-oxohexyl]-5-sulfo-2(3H)-benzoxazolylidene]-1-propenyl]-5-sulfo-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 185397-48-2 HCAPLUS

CN 3H-Indolium, 2-[5-[5-(carboxymethyl)-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-5-[2-[[[2-[3-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-

## MAUPIN 09/829,467

1-(4-sulfobutyl)-3H-indolium-5-yl]methyl]amino]-2-oxoethyl]-3,3-dimethyl-1-(4-sulfobutyl)-, bis(inner salt) (9CI) (CA INDEX NAME)

PAGE 1-B

RN 185397-49-3 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[5-[1-[6-[[2-[3-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-3H-indolium-5-yl]methyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, bis(inner salt) (9CI) (CA INDEX NAME)

RN 185397-50-6 HCAPLUS

CN 3H-Indolium, 5-(aminomethyl)-2-[3-[5-[[[6-[2-[7-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]methyl]-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, bis(inner salt) (9CI) (CA INDEX NAME)

PAGE 1-B

Сн<sub>2</sub>— Nн<sub>2</sub>

RN 185397-51-7 HCAPLUS

CN 3H-Indolium, 1-[6-[[[1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-3H-indolium-5-yl]methyl]amino]-6-oxohexyl]-2-[5-[1-[6-[[1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-3H-indoliumn-5-yl]methyl]amino]-6-oxohexyl]-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, tris(inner salt) (9CI) (CA INDEX NAME)

PAGE 1-B

# IT 146368-10-7 146368-12-9 185397-52-8 185397-53-9 185397-54-0 185397-55-1

RL: PRP (Properties); RCT (Reactant); RACT (Reactant or reagent) (fluorescent labeling complexes with large Stokes shifts prepn. for cell anal.)

RN 146368-10-7 HCAPLUS

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CN 3H-Indolium, 1-(5-carboxypentyl)-2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

RN 146368-12-9 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[7-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

$$Me$$
 $N$ 
 $CH-CH=CH-CH=CH-CH=CH-N$ 
 $N$ 
 $HO_3S$ 
 $Me$ 
 $HO_2C-(CH_2)_5$ 

RN 185397-52-8 HCAPLUS

CN 3H-Indolium, 5-(aminomethyl)-2-[3-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 185397-53-9 HCAPLUS

CN 3H-Indolium, 5-(aminomethyl)-2-[3-[5-(aminomethyl)-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 185397-54-0 HCAPLUS

CN 3H-Indolium, 5-(carboxymethyl)-2-[5-[5-(carboxymethyl)-1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 185397-55-1 HCAPLUS

CN Benzoxazolium, 3-(5-carboxypentyl)-2-[3-[3-(5-carboxypentyl)-5-sulfo-2(3H)-benzoxazolylidene]-1-propenyl]-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

$$CH-CH=CH$$
 $CH-CH=CH$ 
 $CH-CH$ 
 $CH-CH$ 

IT 185397-38-0 185397-40-4 185397-45-9

RL: RCT (Reactant); RACT (Reactant or reagent)
 (fluorescent labeling complexes with large Stokes shifts prepn. for
 cell anal.)

RN 185397-38-0 HCAPLUS

CN 3H-Indolium, 5-amino-2-[5-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$^{-\text{O}_3\text{S}-}$$
 (CH<sub>2</sub>)<sub>4</sub>  $^{-\text{SO}_3\text{H}}$  (CH<sub>2</sub>)<sub>4</sub>  $^{-\text{SO}_3\text{H}}$   $^{-\text{N}_+}$  CH= CH- CH= CH- CH  $^{-\text{N}_+}$  Me Me

RN 185397-40-4 HCAPLUS

CN 3H-Indolium, 5-amino-2-[7-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3,5-heptatrienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

$$^{-O_3S-}$$
 (CH<sub>2</sub>) 4 (CH<sub>2</sub>) 4  $^{-SO_3H}$   $^{N^+}$  CH = CH - CH = CH - CH = Me Me Me

RN 185397-45-9 HCAPLUS

CN 3H-Indolium, 2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-pentadienyl]-1-[6-[(2,5-dioxo-1-pyrrolidinyl)oxy]-6-oxohexyl]-3,3-dimethyl-5-sulfo-, inner salt (9CI) (CA INDEX NAME)

IT 185397-39-1P 185397-46-0DP, succinimidyl ester derivs

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(fluorescent labeling complexes with large Stokes shifts prepn. for cell anal.)

RN 185397-39-1 HCAPLUS

CN 3H-Indolium, 5-[(4,6-dichloro-1,3,5-triazin-2-yl)amino]-2-[5-[1,3-dihydro-3,3-dimethyl-1-(4-sulfobutyl)-2H-indol-2-ylidene]-1,3-pentadienyl]-3,3-dimethyl-1-(4-sulfobutyl)-, inner salt (9CI) (CA INDEX NAME)

RN 185397-46-0 HCAPLUS

CN 3H-Indolium, 1-(5-carboxypentyl)-2-[3-[1-(5-carboxypentyl)-5-[[[6-[2-[5-[1-(5-carboxypentyl)-1,3-dihydro-3,3-dimethyl-5-sulfo-2H-indol-2-ylidene]-1,3-

pentadienyl]-3,3-dimethyl-5-sulfo-3H-indolio]-1-oxohexyl]amino]methyl]-1,3dihydro-3,3-dimethyl-2H-indol-2-ylidene]-1-propenyl]-3,3-dimethyl-5-sulfo, bis(inner salt) (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 1-B

$$= CH - CH = CH$$

$$(CH2)5 - CO2H$$